Barle O'Brysen

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		onyms, and registry numbers, and combine with the concept or neaning. Give examples or relevant citations, authors, etc, if	
known. Please attach a copy of the cover			
Title of Invention: PAG VEVS	time for men	piratory infections disease	شر _و .
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Inventors (please provide full names):	150 90341	naga lace	
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Earliest Priority Filing Date:	12/22/9	<u> </u>	
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Cook 09/868106 Page 1

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L4 72 SEA FILE=REGISTRY ABB=ON C5H9NO4S/MF
L5 38 SEA FILE=REGISTRY ABB=ON L4 NOT RSD/FA
L6 10 SEA FILE=REGISTRY ABB=ON L5 AND (ALANINE OR CYSTEINE)
L7 7 SEA FILE=REGISTRY ABB=ON L6 NOT (ESTER OR N CARBOXYMETHYL)
L9 6 SEA FILE=REGISTRY ABB=ON L7 NOT (N METHOXYCARBONYL)

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L4 72 SEA FILE=REGISTRY ABB=ON C5H9NO4S/MF
L5 38 SEA FILE=REGISTRY ABB=ON L4 NOT RSD/FA
L6 10 SEA FILE=REGISTRY ABB=ON L5 AND (ALANINE OR CYSTEINE)
L7 7 SEA FILE=REGISTRY ABB=ON L6 NOT (ESTER OR N CARBOXYMETHYL)
L9 6 SEA FILE=REGISTRY ABB=ON L7 NOT (N METHOXYCARBONYL)
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Page 2

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545 EA FILE=HCAPLUS ABB=ON L9
         118750 SEA FILE=HCAPLUS ABB=ON RESPIRATORY TRACT+NT/CT
L17
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             49 SEA FILE=HCAPLUS ABB=ON L17 AND L18
L19
              35 SEA FILE=HCAPLUS ABB=ON L19 NOT PY>1997
L26
              72 SEA FILE=REGISTRY ABB=ON C5H9NO4S/MF
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             228 SEA FILE=HCAPLUS ABB=ON
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              49 SEA FILE=HCAPLUS ABB=ON L28 AND P/DT - Patents as document type
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              26 SEA FILE=HCAPLUS ABB=ON L29 AND (INHALANT# OR EXPECTORANT# OR
 L29
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mechanism of
 L31
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56 L26 OR L31 L32

=> fil medl; d que 146 FILE 'MEDLINE' ENTERED AT 09:21:57 ON 09 JUL 2002

FILE LAST UPDATED: 7 JUL 2002 (20020707/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

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252 SEA FILE=MEDLINE ABB=ON CARBOCYSTEINE/CT
          164518 SEA FILE=MEDLINE ABB=ON RESPIRATORY TRACT INFECTIONS+NT/CT
L34
              140 SEA FILE=MEDLINE ABB=ON L34(L)(TU OR PD OR PK OR AD)/CT
L40
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TM - cherapeutie use
PD - pharmacology
PK - pharmacokineties
AD - administration & dosage
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               93 SEA FILE=MEDLINE ABB=ON L43/MAJ
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               26 SEA FILE=MEDLINE ABB=ON L45 NOT PY>1997
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FILE 'MEDLINE' ENTERED AT 09:22:13 ON 09 JUL 2002
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MEDLINE L48 ANSWER 1 OF 81

DUPLICATE 1

Cook 09/868106 Page 3

ACCESSION NUMBER: 82202735 MEDLINE

DOCUMENT NUMBER: 82202735 PubMed ID: 7080939

TITLE: Effect of S-carboxymethylcysteine on the biophysical and

biochemical properties of mucus in chronic bronchitics.

AUTHOR: Cox A; Jabbal-Gill I; Marriott C; Davis S S

SOURCE: ADVANCES IN EXPERIMENTAL MEDICINE AND BIOLOGY, (1982) 144

423-9.

Journal code: 0121103. ISSN: 0065-2598.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198207

ENTRY DATE: Entered STN: 19900317

Last Updated on STN: 20000303 Entered Medline: 19820708

CONTROLLED TERM: Check Tags: Human

*Bronchitis: PP, physiopathology *Carbocysteine: PD, pharmacology

Chronic Disease

*Cysteine: AA, analogs & derivatives

Double-Blind Method

Glycoproteins: ME, metabolism

*Mucus: DE, drug effects Mucus: ME, metabolism Mucus: PH, physiology *Sputum: DE, drug effects Sputum: PH, physiology

Viscosity

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

CHEMICAL NAME: 0 (Glycoproteins)

L48 ANSWER 2 OF 81 MEDLINE

ACCESSION NUMBER: 1998010724 MEDLINE

DOCUMENT NUMBER: 98010724 PubMed ID: 9349882

TITLE: Improvement of mucosal pathology of the sinuses after

exposure to sulfur dioxide by nebulization of

S-carboxymethylcysteine.

AUTHOR: Sugiura Y; Ohashi Y; Nakai Y

CORPORATE SOURCE: Department of Otolaryngology, Osaka City University Medical

School, Japan.

SOURCE: ACTA OTO-LARYNGOLOGICA. SUPPLEMENT, (1997) 531 10-6.

Journal code: 0370355. ISSN: 0365-5237.

PUB. COUNTRY: Norway

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199712

ENTRY DATE: Entered STN: 19980109

Last Updated on STN: 20000303 Entered Medline: 19971209

ABSTRACT:

Since s-carboxymethylcysteine (S-CMC) can directly enhance the ciliary activity in the maxillary sinus mucosa of patients with chronic sinusitis in the absence of significant organic changes of ciliated cells, the nebulization therapy using this medicine might be more effective in the treatment of chronic sinusitis than oral administration of the medicine. The safety of using 0.5-10% of S-SMC as a medicine for nebulization has been experimentally established. The present study was designed to experimentally examine the effectiveness of nebulization using 0.5-10% of S-CMC solution in the treatment of experimental chronic sinusitis in rabbits recurrently exposed to 20 ppm of sulfur dioxide. Thirty-three healthy rabbits were used; 3 of them were used as controls. The remaining 30 were exposed to 20 ppm of sulfur dioxide for 4 h a day for 4

successive weeks. Twelve animals were not treated with any medication during the post-exposure period, and sacrificed at 24 h or 15 days after completion of the final exposure to sulfur dioxide. The remaining 18 animals were treated with nebulization using 10%, 5% or 0.5% of S-CMC solution for 20 min a day for 14 successive days after the final exposure to sulfur dioxide, and they were sacrificed at 24 h after the final nebulization using S-CMC. At the time of sacrifice, the ciliary activity and the morphology of the sinus mucosa were observed to assess the effectiveness of S-CMC nebulization. In the animals sacrificed 24 h after the final exposure, the mucosa of the sinus demonstrated marked epithelial cell injuries, and the ciliary activity was extremely reduced. Complete recovery of the epithelium and the ciliary activity was not recognized in the animals sacrificed 15 days after completion of the exposure. By contrast, epithelial recovery was more accelerated in the animals treated with S-CMC nebulization during the 14 days after the exposure. In the animals treated with 0.5% of S-CMC, the ciliary activity was inferior to that of the control animals, and the epithelial repair was not complete. In the animals treated with 10% of S-CMC, however, ciliary activity and epithelial morphology were completely recovered. In conclusion, our study suggests that clinical application of 10% of S-CMC nebulization may provide otolaryngologists with a new tool in the treatment of sinus diseases such as chronic sinusitis.

Check Tags: Animal CONTROLLED TERM:

*Carbocysteine: TU, therapeutic use

Chronic Disease

Cilia: UL, ultrastructure Epithelium: UL, ultrastructure

Mucociliary Clearance: DE, drug effects

Nasal Mucosa: PA, pathology Nasal Mucosa: UL, ultrastructure

Nebulizers and Vaporizers

Rabbits

Sinusitis: CI, chemically induced

*Sinusitis: DT, drug therapy Sinusitis: PA, pathology Sinusitis: PP, physiopathology

Sulfur Dioxide

2387-59-9 (Carbocysteine); 7446-09-5 (Sulfur Dioxide) CAS REGISTRY NO.:

MEDLINE L48 ANSWER 3 OF 81

1998010723 MEDLINE ACCESSION NUMBER:

PubMed ID: 9349881 98010723 DOCUMENT NUMBER:

Nebulization of S-carboxymethylcysteine does not adversely TITLE: affect the mucociliary system in the paranasal sinus and

trachea of the healthy rabbit.

Sugiura Y; Ohashi Y; Nakai Y AUTHOR:

Department of Otolaryngology, Osaka City University Medical CORPORATE SOURCE:

School, Japan.

ACTA OTO-LARYNGOLOGICA. SUPPLEMENT, (1997) 531 5-9. SOURCE:

Journal code: 0370355. ISSN: 0365-5237.

Norway PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

English LANGUAGE:

Priority Journals FILE SEGMENT:

199712 ENTRY MONTH:

Entered STN: 19980109 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19971209

ABSTRACT:

Chronic sinusitis is a persistent inflammatory impairment of the paranasal sinus. Disturbance of the mucociliary function in the paranasal sinus is the most common finding in chronic sinusitis. S-carboxymethylcysteine (S-CMC) has been shown to directly enhance the ciliary activity of the chronic sinusitis mucosa. Direct contact of the disturbed cilia with S-CMC may recover the

Cook 09/868106 Page 5

reduced beating activity of cilia in chronic sinusitis and the mucosal pathology of the disease can thus be improved. Before S-CMC as medicine for nebulization in the treatment of chronic sinusitis can be clinically applied, however, it should be experimentally established whether nebulization of S-CMC has any adverse effects on the mucociliary system of the respiratory mucosa. The present study was designed to experimentally examine the safety of nebulization of S-CMC especially with regard to the respiratory mucosa. Rabbits were treated with nebulization of three different concentrations of S-CMC solution for 20 min a day for 14 successive days, and their mucosal pathology of the sinus and trachea was examined and compared with that of healthy animals. Nebulization of concentrations of 0.5-10% of S-CMC solution did not affect the ciliary activity in the sinus and tracheal mucosa, nor did this treatment induce pathological changes such as epithelial injury and inflammatory cell accumulation. It is therefore concluded that concentrations of 0.5-10% S-CMC solution are quite safe for the use of nebulization in the treatment of chronic sinusitis.

CONTROLLED TERM: Check Tags: Animal

*Carbocysteine: AD, administration & dosage

Carbocysteine: TU, therapeutic use

Chronic Disease

*Mucociliary Clearance: DE, drug effects

*Nasal Mucosa: DE, drug effects Nasal Mucosa: PH, physiology Nasal Mucosa: UL, ultrastructure

Nebulizers and Vaporizers

*Paranasal Sinuses: DE, drug effects Paranasal Sinuses: PH, physiology Paranasal Sinuses: UL, ultrastructure

Rabbits

Sinusitis: DT, drug therapy Sinusitis: PP, physiopathology

*Trachea: DE, drug effects Trachea: PH, physiology Trachea: UL, ultrastructure 2387-59-9 (Carbocysteine)

CAS REGISTRY NO.:

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L48 ANSWER 4 OF 81 MEDLINE

ACCESSION NUMBER: 93273289 MEDLINE

DOCUMENT NUMBER: 93273289 PubMed ID: 8500784

TITLE: Carbocisteine improves the mucociliary transport rate in

rats with SO2-induced bronchitis.

AUTHOR: Zahm J M; Levrier J; Duval D; Pierrot D; Puchelle E

CORPORATE SOURCE: INSERM U 314, CHR Maison Blanche, Reims, France.

SOURCE: FUNDAMENTAL AND CLINICAL PHARMACOLOGY, (1993) 7 (3-4)

155-60.

Journal code: 8710411. ISSN: 0767-3981.

PUB. COUNTRY: DOCUMENT TYPE:

France

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199306

ENTRY DATE:

Entered STN: 19930716

Last Updated on STN: 19930716 Entered Medline: 19930630

ABSTRACT:

In order to study the effect of carbocisteine on the mucociliary function of the respiratory tract, we performed a double-blind study on rats with SO2-induced (400 ppm) hypersecretion. During the experimental bronchitis, the treated group of rats received carbocisteine through a stomach tube at a dose level of 500 mg/kg for 15 days, whereas the untreated group of rats received distilled water. After killing the rats, and following lung excision, the respiratory mucus was scraped off and collected by using a glass capillary. The

mucus degree of purulence was macroscopically estimated and the mucus transport rate was measured by using the frog palate technique. The mean mucus relative transport rate, measured on the frog palate, was 0.60 ± -0.17 in the untreated group and was significantly higher (P < 0.01) in the treated group (0.73 +/-0.14). Carbocisteine also significantly altered (P < 0.01) the mucus macroscopical aspect, leading to a decrease in the number of rats with purulent mucus. These results suggest that carbocisteine maintains an efficient mucus transport rate, leading to a less infected respiratory tract.

Check Tags: Animal; Male CONTROLLED TERM:

*Bronchitis: PP, physiopathology *Carbocysteine: PD, pharmacology

Double-Blind Method Microscopy, Electron

*Mucociliary Clearance: DE, drug effects Mucous Membrane: UL, ultrastructure

Mucus: ME, metabolism

Rats

Rats, Sprague-Dawley

Respiratory System: UL, ultrastructure

Sulfur Dioxide

2387-59-9 (Carbocysteine); 7446-09-5 (Sulfur Dioxide) CAS REGISTRY NO .:

MEDLINE L48 ANSWER 5 OF 81

94078090 MEDLINE ACCESSION NUMBER:

PubMed ID: 8256077 94078090 DOCUMENT NUMBER:

Effect of S-carboxymethylcysteine on ciliary activity in TITLE:

chronic sinusitis.

Ohashi Y; Nakai Y; Sugiura Y; Ohno Y; Okamoto H; Hayashi M AUTHOR:

Department of Otolaryngology, Osaka City University Medical CORPORATE SOURCE:

School, Japan.

RHINOLOGY, (1993 Sep) 31 (3) 107-11. SOURCE: Journal code: 0347242. ISSN: 0300-0729.

Netherlands PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

English LANGUAGE:

Priority Journals FILE SEGMENT:

199401 ENTRY MONTH:

Entered STN: 19940203 ENTRY DATE:

Last Updated on STN: 19940203 Entered Medline: 19940111

This study was designed to investigate the possible pharmacological effect of S-carboxy-methylcysteine (S-CMC) on the ciliary activity, using an in vitro experimental system after removing mucus. Ciliary activity from healthy rabbit maxillary sinus and from healthy human nasal mucosa demonstrated no significant change in RPMI 1640 containing S-CMC. On the other hand, the effect of S-CMC on the reduced ciliary activity from patients with chronic sinusitis was quite varied among the cases examined. S-CMC demonstrated no stimulatory effect on the beating activity of cilia that have a baseline activity of less than 400 beats/min. However, S-CMC was able to enhance the beating activity of cilia that demonstrated a baseline activity of more than 400 beats/min. S-CMC at 0.5% induced a larger ciliostimulatory effect than 0.05% S-CMC. In conclusion, our study has clearly demonstrated that S-CMC could directly enhance ciliary activity of chronic sinusitis in the absence of significant organic change of ciliated cells.

Check Tags: Animal; Human; In Vitro CONTROLLED TERM: *Carbocysteine: PD, pharmacology

Chronic Disease

Cilia: DE, drug effects Cilia: PH, physiology

Maxillary Sinus: DE, drug effects

Cook 09/868106 Page 7

*Maxillary Sinus: PP, physiopathology

*Maxillary Sinusitis: PP, physiopathology

Nasal Mucosa: PH, physiology

Rabbits

2387-59-9 (Carbocysteine) CAS REGISTRY NO.:

L48 ANSWER 6 OF 81 MEDITNE

93008414 ACCESSION NUMBER: MEDIT NE

93008414 DOCUMENT NUMBER: PubMed ID: 1394568

[Carbocysteine in the treatment of recurrent bronchitis in TITLE:

infants].

Karbocystein v liecbe recidivujucich bronchitid u dojciat.

Banovcin P; Jakusova L; Rosslerova V; Miklerova M; Pullmann AUTHOR:

Detska klinika Jeseniovej lekarskej fakulty Univerzity CORPORATE SOURCE:

Komenskeho, Martin.

CESKOSLOVENSKA PEDIATRIE, (1992 Sep) 47 (9) 543-6. SOURCE:

Journal code: 0403576. ISSN: 0069-2328.

Czechoslovakia PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: Slovak

Priority Journals FILE SEGMENT:

ENTRY MONTH: 199211

ENTRY DATE: Entered STN: 19930122

> Last Updated on STN: 19930122 Entered Medline: 19921125

ABSTRACT:

In a group of 51 children aged 6-24 months the therapeutic effectiveness of the mucolytic preparation carbocysteine was tested and compared with the effect of Ipeca syrup. The effect was evaluated by means of a point score comprising changes of the clinical picture of the disease and the use of other laboratory examinations. The results of the examination revealed the more favourable effect of carbocysteine, as compared with a mixture of Ipeca syrup in the treatment of acute relapsing bronchitis in infants.

CONTROLLED TERM: Check Tags: Female; Human; Male

> *Bronchitis: DT, drug therapy *Carbocysteine: TU, therapeutic use

Child, Preschool English Abstract

Infant Recurrence

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine)

L48 ANSWER 7 OF 81 MEDLINE

ACCESSION NUMBER: 93138542 MEDLINE

93138542 PubMed ID: 1487227 DOCUMENT NUMBER:

Study on the effect of oral administration of carbocysteine TITLE:

on ventilatory parameters in the SO2 inhalation model of

bronchitis in the rat.

AUTHOR: Levrier J; Duval D; Lloyd K G

CORPORATE SOURCE: Synthelabo Recherche, (LERS) Biology Department, Bagneux,

France.

SOURCE: FUNDAMENTAL AND CLINICAL PHARMACOLOGY, (1992) 6 (6) 231-6.

Journal code: 8710411. ISSN: 0767-3981.

PUB. COUNTRY: France

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199302

Entered STN: 19930312 ENTRY DATE:

> Last Updated on STN: 19930312 Entered Medline: 19930224

ABSTRACT: In order to study the physiological correlates of the beneficial action of carbocisteine (S-carboxy-methyl-cysteine), we have measured the changes occurring in ventilatory parameters in rats made bronchitic by prolonged exposure (2 weeks) to air containing sulfur dioxide (SO2). In animals treated with distilled water (1 ml/100 g/day), statistically significant (P < 0.05) changes in respiratory frequency (-20%) and tidal volume (+31%) were found. As a result of these opposing changes, the ventilation/min was stable. Moreover, the compliance was decreased (33%, P < 0.05) and the resistance was greatly enhanced (+ 99%, P < 0.05). The concomitant administration of carbocisteine (500 mg/kg po/day) with SO2 inhalation significantly (P < 0.05) prevented the development of resistance without effecting significant changes in the other parameters except for a slight improvement in ventilation/min. In conclusion, this improved respiratory resistance in the bronchitic carbocisteine-treated animals tallies with a decrease in mucus retention associated with the return to normal of rheological characteristics of the secreted mucus.

Check Tags: Animal; Male CONTROLLED TERM:

Administration, Inhalation

Administration, Oral

Bronchitis: CI, chemically induced

*Bronchitis: DT, drug therapy Bronchitis: PP, physiopathology *Carbocysteine: TU, therapeutic use

Disease Models, Animal Lung: DE, drug effects Random Allocation

Rats

Rats, Sprague-Dawley

*Respiration: DE, drug effects Respiration: PH, physiology Respiratory Function Tests

Sulfur Dioxide

2387-59-9 (Carbocysteine); 7446-09-5 (Sulfur Dioxide) CAS REGISTRY NO .:

MEDLINE L48 ANSWER 8 OF 81

MEDLINE 92210058 ACCESSION NUMBER:

PubMed ID: 1555809 92210058 DOCUMENT NUMBER:

Effects of S-carboxymethyl-L-cysteine on pulmonary sialyl TITLE:

transferase activity in vitro, in healthy and in

sulphur-dioxide-induced bronchitic rats.

Berry C N; Lloyd K G; Louisot P AUTHOR:

Synthelabo Recherche (LERS), Bagneux, France. CORPORATE SOURCE:

FUNDAMENTAL AND CLINICAL PHARMACOLOGY, (1992) 6 (1) 29-35. SOURCE:

Journal code: 8710411. ISSN: 0767-3981.

France PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

English LANGUAGE:

Priority Journals FILE SEGMENT:

199205 ENTRY MONTH:

Entered STN: 19920515 ENTRY DATE:

Last Updated on STN: 19980206 Entered Medline: 19920501

S-carboxymethyl-L-cysteine (carbocysteine) improves the visco-elastic properties of bronchial mucus in vivo, possibly as a result of an increase in the relative proportions of sialomucins in bronchial mucus. Carbocysteine was therefore studied in vitro and ex vivo in both normal and bronchitic rats on pulmonary sialyl transferase, responsible for the addition of sialic acid to mucus glycoproteins. Bronchitis was induced in male Sprague-Dawley rats by repeated exposure to sulphur dioxide for two weeks. During this time they received either 500 mg kg-1 day-1 carbocysteine or its vehicle by the oral route. Rats not being exposed to SO2 received the same treatment. The animals Cook 09/868106 Page 9

were then killed, and subcellular fractions prepared by differential centrifugation of lung homogenates. Sialyl transferase was assayed using CMP-14C sialic acid as substrate and desialysed fetuin as exogenous acceptor. Enzyme activity was located in both the (Golgi-containing) 10,000 g and 100,000 g pellets with minor activity in the cytosolic supernatants. When tested in vitro between 10(-6) and 10(-3) M, carbocysteine had no effect on sialyl transferase activity in microsomes taken from healthy or bronchitis rats. Repeated administration of carbocysteine was without effect on the sialyl transferase activity in 10,000 g pellets taken from healthy rats. However, in bronchitic rats there was a small but statistically significant (P less than 0.05) increase in enzymic activity in the treated group compared to the animals receiving the vehicle. There was no difference in the activity of the microsomal enzyme compared to vehicle-treated controls in either healthy or bronchitic rats. We conclude that it is possible that an increase in sialyl transferase activity in a Golgi-containing fraction of bronchitic lungs could explain the relative increase in sialomucins in bronchitic subjects.

CONTROLLED TERM: Check Tags: Animal; Male

Bronchitis: CI, chemically induced

*Bronchitis: EN, enzymology

Carbocysteine: AD, administration & dosage

*Carbocysteine: PD, pharmacology

*Lung: EN, enzymology

Rats

Rats, Inbred Strains

*Sialyltransferases: AN, analysis Subcellular Fractions: EN, enzymology

Sulfur Dioxide

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 7446-09-5 (Sulfur Dioxide)

CHEMICAL NAME: EC 2.4.99.- (Sialyltransferases)

L48 ANSWER 9 OF 81 MEDLINE

ACCESSION NUMBER: 91288942 MEDLINE

DOCUMENT NUMBER: 91288942 PubMed ID: 2099568

TITLE: Long-lasting effects on rheology and clearance of bronchial

mucus after short-term administration of high doses of carbocysteine-lysine to patients with chronic bronchitis.

AUTHOR: Braga P C; Allegra L; Rampoldi C; Ornaghi A; Beghi G

CORPORATE SOURCE: Center for Respiratory Pharmacology, School of Medicine,

University of Milan, Italy.

SOURCE: RESPIRATION, (1990) 57 (6) 353-8.

Journal code: 0137356. ISSN: 0025-7931.

PUB. COUNTRY: Switzerland

DOCUMENT TYPE: (CLINICAL TRIAL)

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199108

ENTRY DATE: Entered STN: 19910825

Last Updated on STN: 19960129

Entered Medline: 19910805

ABSTRACT:

The rheological behavior and clearance of bronchial mucus samples collected by protected expectoration from 24 out-patients with simple chronic bronchitis were investigated before, at the end of a short period of treatment (4 days) with a single oral dose of 2.7 g (sachet) of carbocysteine-lysine (evening meal), and on the 4th and 8th days after the end of treatment versus placebo. In the group treated with carbocysteine-lysine, there were significant reductions in viscosity (-67, -48, -62%) and increases in mucociliary transport (+41, +31, +34%) at the three times mentioned. The most striking finding was that the improvements were still present 8 days after cessation of treatment. The elasticity parameter was not affected in any statistically significant way

Page 10 09/868106 Cook

(-10, -24, +65%). These findings suggest the presence of some type of 'post-mucoactive' effect.

Check Tags: Female; Human; Male CONTROLLED TERM:

Adult Aged

*Bronchitis: ME, metabolism

*Carbocysteine: PK, pharmacokinetics

Chronic Disease Middle Age

Mucociliary Clearance *Mucus: ME, metabolism Random Allocation

Rheology Viscosity

2387-59-9 (Carbocysteine) CAS REGISTRY NO.:

MEDLINE L48 ANSWER 10 OF 81

MEDLINE 91315222 ACCESSION NUMBER:

PubMed ID: 3155012 91315222 DOCUMENT NUMBER:

TITLE:

[Carbocysteine-sobrerol combination and exacerbation of

chronic bronchitis].

Associazione carbocisteina-sobrerolo e riacutizzazioni

della bronchite cronica.

Pasturenzi L; Donnetta A M; Gualtieri G; Luisetti M AUTHOR:

ARCHIVIO MONALDI PER LE MALATTIE DEL TORACE, (1988 Nov-Dec) SOURCE:

43 (6) 487-505. Ref: 45

Journal code: 8902999. ISSN: 1120-0391.

Italy PUB. COUNTRY:

(CLINICAL TRIAL) DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE:

Italian

Priority Journals FILE SEGMENT:

199108 ENTRY MONTH:

Entered STN: 19910913 ENTRY DATE:

Last Updated on STN: 19910913 Entered Medline: 19910829

CONTROLLED TERM:

Check Tags: Comparative Study; Human

Ambroxol: TU, therapeutic use

Amoxicillin: AD, administration & dosage

Amoxicillin: TU, therapeutic use *Bronchitis: DT, drug therapy

*Carbocysteine: AD, administration & dosage

Cefuroxime: AD, administration & dosage

Cefuroxime: TU, therapeutic use

Chronic Disease

Drug Therapy, Combination

English Abstract

*Expectorants: AD, administration & dosage *Terpenes: AD, administration & dosage

Time Factors

18683-91-5 (Ambroxol); 2387-59-9 (Carbocysteine); CAS REGISTRY NO.:

26787-78-0 (Amoxicillin); 498-71-5 (sobrerol); 55268-75-2

(Cefuroxime)

0 (Expectorants); 0 (Terpenes) CHEMICAL NAME:

MEDLINE L48 ANSWER 11 OF 81

MEDLINE 89100485 ACCESSION NUMBER:

PubMed ID: 3062806 89100485

DOCUMENT NUMBER:

[Comparative evaluation of the effectiveness of lasolvan and mucodine in chronic nonspecific lung diseases]. TITLE:

Cook 09/868106 Page 11

Sravnitel'naia otsenka effektivnosti lasol'vana i mukodina

pri khronicheskikh nespetsificheskikh zabolevaniiakh

legkikh.

AUTHOR: Solopov V N; Kolganova N A

SOURCE: SOVETSKAIA MEDITSINA, (1988) (5) 69-72.

Journal code: 0404525. ISSN: 0038-5077.

PUB. COUNTRY: USSR

DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Russian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198902

ENTRY DATE: Entered STN: 19900308

Last Updated on STN: 20000303 Entered Medline: 19890222

CONTROLLED TERM: Check Tags: Comparative Study; Human

*Ambroxol: TU, therapeutic use

*Asthma: DT, drug therapy

*Bromhexine: AA, analogs & derivatives

*Bronchitis: DT, drug therapy

*Carbocysteine: TU, therapeutic use

Chronic Disease Clinical Trials

*Cysteine: AA, analogs & derivatives

CAS REGISTRY NO.: 18683-91-5 (Ambroxol); 2387-59-9 (Carbocysteine); 3572-43-8

(Bromhexine); 52-90-4 (Cysteine)

L48 ANSWER 12 OF 81 MEDLINE

ACCESSION NUMBER: 86062057 MEDLINE

DOCUMENT NUMBER: 86062057 PubMed ID: 4067726

TITLE: Effects of carbocysteine on experimental chronic sinusitis

caused by long-term exposure to SO2.

AUTHOR: Ohashi Y; Nakai Y; Koshimo H; Ikeoka H; Maruoka K; Takagi K

SOURCE: NIPPON JIBIINKOKA GAKKAI KAIHO [JOURNAL OF THE

OTO-RHINO-LARYNGOLOGICAL SOCIETY OF JAPAN], (1985 Aug) 88

(8) 1056-60.

Journal code: 7505728. ISSN: 0030-6622.

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Japanese

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198601

ENTRY DATE: Entered STN: 19900321

Last Updated on STN: 20000303 Entered Medline: 19860114

CONTROLLED TERM: Check Tags: Animal

*Carbocysteine: TU, therapeutic use

Chronic Disease

*Cysteine: AA, analogs & derivatives

English Abstract

*Maxillary Sinus: UL, ultrastructure

Microscopy, Electron

Rabbits

Sinusitis: CI, chemically induced

*Sinusitis: PA, pathology
*Sulfur Dioxide: TO, toxicity

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine); 7446-09-5

(Sulfur Dioxide)

L48 ANSWER 13 OF 81 MEDLINE

ACCESSION NUMBER: 86077525 MEDLINE

DOCUMENT NUMBER: 86077525 PubMed ID: 3907681

TITLE: Long-term oral carbocisteine therapy in patients with

chronic bronchitis. A double blind trial with placebo

control.

Grillage M; Barnard-Jones K

BRITISH JOURNAL OF CLINICAL PRACTICE, (1985 Oct) 39 (10) AUTHOR: SOURCE:

395-8.

Journal code: 0372546. ISSN: 0007-0947.

ENGLAND: United Kingdom PUB. COUNTRY:

(CLINICAL TRIAL) DOCUMENT TYPE:

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

198602 ENTRY MONTH:

. Entered STN: 19900321 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19860212

Check Tags: Human CONTROLLED TERM:

Adult

*Bronchitis: DT, drug therapy Bronchitis: PP, physiopathology Carbocysteine: AE, adverse effects *Carbocysteine: TU, therapeutic use

Clinical Trials

*Cysteine: AA, analogs & derivatives

Double-Blind Method Peak Expiratory Flow Rate

2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO.:

MEDLINE L48 ANSWER 14 OF 81

MEDLINE 85305322

ACCESSION NUMBER: PubMed ID: 4037622 85305322

DOCUMENT NUMBER: [Changes in IgA levels in nasal mucus after upper TITLE:

respiratory tract diseases in infants treated with

carbocysteine].

Modifications du taux des IgA du mucus nasal au decours des affections des voies aeriennes superieures du nourrisson

traitees par la carbocisteine.

Henocq A; Moreau C; Mallet E; Sauger F; de Menibus C H ANNALES D OTO-LARYNGOLOGIE ET DE CHIRURGIE CERVICO-FACIALE, AUTHOR: SOURCE:

(1985) 102 (5) 373-5.

Journal code: 9431026. ISSN: 0003-438X.

France PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

French LANGUAGE:

Priority Journals FILE SEGMENT:

198510 ENTRY MONTH:

Entered STN: 19900320 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19851007

ABSTRACT:

The authors have studied IgA level in nasal mucus of children, either not treated-controls, or treated with carbocysteine. All had common rhinobronchial diseases. They have noted a significant increase in IgA level in the treated group, from the 7th day.

Check Tags: Human CONTROLLED TERM:

*Carbocysteine: TU, therapeutic use

Child, Preschool

*Cysteine: AA, analogs & derivatives

English Abstract

*Immunoglobulin A, Secretory: AN, analysis

Infant

*Nasal Mucosa: IM, immunology

Cook 09/868106 Page 13

*Respiratory Tract Infections: DT, drug therapy Respiratory Tract Infections: IM, immunology

Time Factors

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

CHEMICAL NAME: 0 (Immunoglobulin A, Secretory)

L48 ANSWER 15 OF 81 MEDLINE

ACCESSION NUMBER: 86058159 MEDLINE

DOCUMENT NUMBER: 86058159 PubMed ID: 4066083

TITLE: Comparison between penetration of amoxicillin combined with

carbocysteine and amoxicillin alone in pathological

bronchial secretions and pulmonary tissue.

AUTHOR: Braga P C; Scaglione F; Scarpazza G; Fraticelli G; Roviaro

G; Varoli F; Mariani C; Falchi M; Fraschini F

SOURCE: INTERNATIONAL JOURNAL OF CLINICAL PHARMACOLOGY RESEARCH,

(1985) 5 (5) 331-40.

Journal code: 8110183. ISSN: 0251-1649.

PUB. COUNTRY: Switzerland

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198601

ENTRY DATE: Entered STN: 19900321

Last Updated on STN: 20000303 Entered Medline: 19860108

ABSTRACT:

Patients with chronic bronchitis were treated orally with either amoxicillin (500 mg) alone or in combination with carbocysteine (150 mg), thrice daily for five days, in order to assess whether the combination allows higher antibiotic levels to be obtained in bronchial mucus than those obtained from amoxicillin alone. Serum and mucus levels were determined for each patient at first and fifth day of the two drug regimens. The levels of amoxicillin in the lung tissue collected in patients undergoing pulmonary surgery were also determined after a single oral dose of amoxicillin (1 g) or of amoxicillin (1 g) plus carbocysteine (300 mg). In the bronchial secretions, at the same plasma concentrations, amoxicillin levels were statistically higher after administration of combined substances. These findings indicate the presence of a pharmacokinetic synergism between these compounds, which allows amoxicillin to penetrate more easily through the hemato-bronchial barrier. The association of amoxicillin and carbocysteine, determining an increase of the quantitative levels of antibiotic in the bronchial secretion (also if it is purulent), performs a sterilizing action in a short time with significant therapeutic advantages.

CONTROLLED TERM: Check Tags: Female; Human; Male

Aged

Amoxicillin: AD, administration & dosage

*Amoxicillin: TU, therapeutic use

Bronchi: BS, blood supply *Bronchi: SE, secretion

*Bronchitis: DT, drug therapy Bronchitis: MI, microbiology Bronchitis: PA, pathology

Carbocysteine: AD, administration & dosage

*Carbocysteine: TU, therapeutic use *Cysteine: AA, analogs & derivatives

Drug Interactions

Drug Therapy, Combination

*Lung: PA, pathology

Middle Age

Mucus: SE, secretion

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 26787-78-0 (Amoxicillin);

52-90-4 (Cysteine)

MEDLINE L48 ANSWER 16 OF 81

MEDLINE 86268238 ACCESSION NUMBER:

PubMed ID: 3836611 86268238 DOCUMENT NUMBER:

TITLE:

[Effect of S-carboxymethylcysteine on the concentration of antibiotics in bronchial secretions and its therapeutic

effects].

Studio dell'attivita della S-carbossimetilcisteina sulle concentrazioni di antibiotici nel secreto bronchiale ed

effetti terapeutici.

Pirali F; Ravizzola G; Santus G; Inzoli M R; Turano A AUTHOR: ARCHIVIO MONALDI PER LA TISIOLOGIA E LE MALATTIE DELL APPARATO RESPIRATORIO, (1985 Jan-Apr) 40 (1-2) 3-18. Journal code: 1263173. ISSN: 0004-0185. SOURCE:

Italy PUB. COUNTRY:

(CLINICAL TRIAL) DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

Italian LANGUAGE:

Priority Journals FILE SEGMENT:

198608 ENTRY MONTH:

Entered STN: 19900321 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19860821

Check Tags: Female; Human; Male CONTROLLED TERM:

Aged

Antibiotics: ME, metabolism

Bronchopneumonia: DT, drug therapy *Bronchopneumonia: ME, metabolism *Carbocysteine: PD, pharmacology *Cysteine: AA, analogs & derivatives

Drug Therapy, Combination

English Abstract

Middle Age

Sputum: ME, metabolism

2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO.:

0 (Antibiotics) CHEMICAL NAME:

MEDLINE L48 ANSWER 17 OF 81

81272536 MEDLINE ACCESSION NUMBER:

PubMed ID: 7022385 81272536 DOCUMENT NUMBER:

[Mucodine in the treatment of chronic bronchitis]. TITLE:

Zastosowanie mukodyny w leczeniu przewleklego zapalenia

oskrzeli.

Wierzbicka M; Wojcik R A AUTHOR:

PNEUMONOLOGIA POLSKA, (1981) 49 (5) 369-76. SOURCE:

Journal code: 7605692. ISSN: 0376-4761.

Poland PUB. COUNTRY:

(CLINICAL TRIAL) DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

Polish LANGUAGE:

Priority Journals FILE SEGMENT:

198110 ENTRY MONTH:

Entered STN: 19900316 ENTRY DATE:

Last Updated on STN: 20000303 Entered Medline: 19811029

Check Tags: Female; Human; Male CONTROLLED TERM:

Adolescence

Adult

*Bronchitis: DT, drug therapy *Carbocysteine: TU, therapeutic use

Chronic Disease

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Clinical Trials

*Cysteine: AA, analogs & derivatives

Double-Blind Method English Abstract

Middle Age Placebos

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

CHEMICAL NAME: 0 (Placebos)

L48 ANSWER 18 OF 81 MEDLINE

ACCESSION NUMBER: 81237546 MEDLINE

DOCUMENT NUMBER: 81237546 PubMed ID: 7250579

DOCUMENT NOTIFIES.

TITLE: [Absorption, elimination and therapeutic effectiveness of a

new antibiotic and mucolytic combination for oral

administration].

Studio sull'assorbimento, sull'eliminazione e sulla

efficacia clinica di una nuova associazione

antibiotico-mucolitica per via orale.

AUTHOR: Silvia G; Giambrone F; Battaglia E; Romano M

SOURCE: GIORNALE DI CLINICA MEDICA, (1981 Mar) 62 (3) 209-27.

Journal code: 0413411. ISSN: 0017-0275.

PUB. COUNTRY: Italy

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Italian

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198109

ENTRY DATE: Entered STN: 19900316

Last Updated on STN: 20000303 Entered Medline: 19810922

CONTROLLED TERM: Check Tags: Female; Human; Male

Adult Aged

*Bacterial Infections: DT, drug therapy

Bronchitis: DT, drug therapy Bronchopneumonia: DT, drug therapy

Carbocysteine: AD, administration & dosage

Carbocysteine: ME, metabolism

*Carbocysteine: TU, therapeutic use

Cefadroxil

Cephalexin: AD, administration & dosage *Cephalexin: AA, analogs & derivatives

Cephalexin: ME, metabolism
Cephalexin: TU, therapeutic use
*Cysteine: AA, analogs & derivatives

Drug Therapy, Combination

English Abstract

Middle Age

*Respiratory Tract Infections: DT, drug therapy

CAS REGISTRY NO.: 15686-71-2 (Cephalexin); 2387-59-9 (Carbocysteine);

50370-12-2 (Cefadroxil); 52-90-4 (Cysteine)

L48 ANSWER 19 OF 81 MEDLINE

ACCESSION NUMBER: 81177684 MEDLINE

DOCUMENT NUMBER: 81177684 PubMed ID: 7013137

TITLE: [Optimal use of expectorants (current trends)].

Optimal'noe primenenie otkharkivaiushchikh preparatov

(sovremennye tendentsii).

AUTHOR: Mirrakhimov M M; Brimkulov N N; Rafibekova Zh S

SOURCE: TERAPEVTICHESKII ARKHIV, (1981) 53 (1) 110-7. Ref: 114

Journal code: 2984818R. ISSN: 0040-3660.

PUB. COUNTRY: USSF

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

LANGUAGE:

Russian

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198106

ENTRY DATE:

Entered STN: 19900316

Last Updated on STN: 20000303 Entered Medline: 19810613

CONTROLLED TERM:

Check Tags: Human; In Vitro

Biological Transport

*Bromhexine: TU, therapeutic use

Bronchi: SE, secretion

*Bronchitis: DT, drug therapy *Carbocysteine: TU, therapeutic use

Chronic Disease

*Cysteine: AA, analogs & derivatives

Elasticity

Sputum: DE, drug effects Sputum: ME, metabolism

Viscosity

CAS REGISTRY NO .:

2387-59-9 (Carbocysteine); 3572-43-8 (Bromhexine); 52-90-4

(Cysteine)

L48 ANSWER 20 OF 81

MEDLINE

ACCESSION NUMBER: DOCUMENT NUMBER:

MEDLINE 79221195

TITLE:

PubMed ID: 460097 79221195

[Changes in sputum in catarrhal bronchitis in children after treatment with S-carboxymethylcysteine (viscosimetric

studies)].

Modificazioni dell'escreato nella bronchite catarrale in

eta pediatrica dopo trattamento con S-

carbossimetilcisteina. (Indagine viscosimetrica).

AUTHOR: SOURCE: Castello D; Costa G; De Candussio G

MINERVA PEDIATRICA, (1979 Mar 15) 31 (5) 371-80.

Journal code: 0400740. ISSN: 0026-4946.

PUB. COUNTRY:

Italy

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

Italian

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

197909

ENTRY DATE:

Entered STN: 19900315 Last Updated on STN: 19900315

Entered Medline: 19790925

CONTROLLED TERM:

Check Tags: Female; Human; Male

Administration, Oral

*Bronchitis: DT, drug therapy

Carbocysteine: AD, administration & dosage

*Carbocysteine: TU, therapeutic use

Child

Child, Preschool

*Cysteine: AA, analogs & derivatives

Drug Evaluation English Abstract

Infant Viscosity

2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO.:

L48 ANSWER 21 OF 81 MEDLINE

ACCESSION NUMBER:

MEDLINE

79106561 PubMed ID: 367726 79106561

DOCUMENT NUMBER: TITLE:

Effects of S-carboxymethylcysteine on tracheal mucus

velocity.

AUTHOR:

Goodman R M; Yergin B M; Sackner M A

CHEST, (1978 Dec) 74 (6) 615-8. SOURCE:

Journal code: 0231335. ISSN: 0012-3692.

Cook 09/868106 Page 17

PUB. COUNTRY: DOCUMENT TYPE: United States

(CLINICAL TRIAL)

(CONTROLLED CLINICAL TRIAL) Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 197904

ENTRY DATE:

Entered STN: 19900315

Last Updated on STN: 19980206 Entered Medline: 19790425

ABSTRACT:

The effects of S-carboxymethylcysteine on tracheal mucus velocity were assessed in a double blind crossover study between 2 grams S-carboxymethylcysteine and placebo. Subjects included six healthy non-smokers, eight smokers with small airway disease and chronic simple bronchitis, and eight subjects with chronic obstructive bronchitis. Tracheal mucus velocity was measured prior to and two and three hours after each subject had ingested S-carboxymethylcysteine or placebo. No significant change in tracheal mucus velocity occurred after placebo or S-carboxymethylcysteine in any of the groups, indicating that the drug has no acute effect on mucus transport.

CONTROLLED TERM:

Check Tags: Female; Human; Male; Support, U.S. Gov't,

P.H.S. Adult

> Bronchitis: DT, drug therapy *Carbocysteine: PD, pharmacology Carbocysteine: TU, therapeutic use

Chronic Disease Clinical Trials

*Cysteine: AA, analogs & derivatives

Lung Diseases, Obstructive: DT, drug therapy

Middle Age

*Mucus: DE, drug effects

Smoking

*Trachea: DE, drug effects

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

L48 ANSWER 22 OF 81 MEDLINE

79085799 MEDLINE ACCESSION NUMBER:

79085799 PubMed ID: 365537 DOCUMENT NUMBER:

Effect of the mucoregulator S-carboxy-methyl-cysteine in TITLE:

patients with chronic bronchitis.

AUTHOR: SOURCE:

Puchelle E; Aug F; Polu J M

EUROPEAN JOURNAL OF CLINICAL PHARMACOLOGY, (1978 Nov 27) 14

(3) 177-84.

Journal code: 1256165. ISSN: 0031-6970.

PUB. COUNTRY:

GERMANY, WEST: Germany, Federal Republic of

DOCUMENT TYPE:

(CLINICAL TRIAL)

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

197903

ENTRY DATE:

Entered STN: 19900315

Last Updated on STN: 19900315 Entered Medline: 19790313

ABSTRACT:

Twenty patients with stable chronic bronchitis entered a double-blind study in which changes in clinical and respiratory function and biochemical and rheological variations were examined after treatment with the mucoregulator S-carboxy-methyl-cysteine (S.C.M.C.). After one week of single-blind placebo, a two week double-blind study was initiated with placebo or oral S.C.M.C. 3 q/24h. After two weeks, a significant clinical improvement was observed in

patients treated with S.C.M.C. During treatment, there was no change in respiratory function, although a drop in FEV1/VC was noted in the placebo group. A significant increase in the viscosity of the expectorations was observed after treatment with S.C.M.C. for two weeks. The levels of secretory IgA and of serum albumin in the expectorations remained stable, whereas in the placebo group, there was a slight but significant increase in serum albumin. In this group of non-infected chronic bronchitic patients, S.C.M.C. appeared to normalize the secretory function of the bronchial mucosa by preventing inflammation and enhancing the viscoelastic properties of bronchial secretions.

Check Tags: Human; Male CONTROLLED TERM: Aged

*Bronchitis: DT, drug therapy Bronchitis: MI, microbiology Bronchitis: PP, physiopathology Carbocysteine: AE, adverse effects *Carbocysteine: TU, therapeutic use

Chronic Disease Clinical Trials

*Cysteine: AA, analogs & derivatives

Double-Blind Method

Middle Age Placebos

Respiratory Function Tests Sputum: AN, analysis Sputum: DE, drug effects Sputum: MI, microbiology

2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO .:

0 (Placebos) CHEMICAL NAME:

MEDLINE L48 ANSWER 23 OF 81

MEDLINE 77107156 ACCESSION NUMBER:

PubMed ID: 797159 77107156

DOCUMENT NUMBER:

[The treatment of bronchitic syndrome using Transbronchin TITLE: in the practice].

Die Behandlung des bronchitischen Syndroms mit

Transbronchin in der Praxis.

ZFA. ZEITSCHRIFT FUR ALLGEMEINMEDIZIN, (1976 Dec 20) 52 AUTHOR: SOURCE:

(35) 1832-4.

Journal code: 7613263. ISSN: 0341-9835. GERMANY, WEST: Germany, Federal Republic of

PUB. COUNTRY: (CLINICAL TRIAL) DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

German LANGUAGE:

Priority Journals FILE SEGMENT:

197703

ENTRY MONTH: Entered STN: 19900313 ENTRY DATE:

Last Updated on STN: 19900313 Entered Medline: 19770321

Check Tags: Female; Human; Male CONTROLLED TERM:

Adolescence

Adult

Aged *Bronchitis: DT, drug therapy *Carbocysteine: TU, therapeutic use

Clinical Trials

*Cysteine: AA, analogs & derivatives

Middle Age

Syndrome 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO .:

MEDLINE L48 ANSWER 24 OF 81

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77025332 ACCESSION NUMBER: MEDLINE

PubMed ID: 789027 DOCUMENT NUMBER: 77025332

TITLE: S-carboxymethylcysteine in the fluidification of sputum and

treatment of chronic airway obstruction.

AUTHOR: Edwards G F; Steel A E; Scott J K; Jordan J W

CHEST, (1976 Oct) 70 (4) 506-13. SOURCE:

Journal code: 0231335. ISSN: 0012-3692.

PUB. COUNTRY: United States (CLINICAL TRIAL) DOCUMENT TYPE:

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

Abridged Index Medicus Journals; Priority Journals FILE SEGMENT:

197612 ENTRY MONTH:

Entered STN: 19900313 ENTRY DATE:

> Last Updated on STN: 19980206 Entered Medline: 19761223

ABSTRACT:

The clinical results and changes in sputum found in both a short-term inpatient trial and a subsequent long-term outpatient investigation (three-month double-blind controlled study) of 82 patients with chronic bronchitis treated with a new mucolytic agent, S-carboxymethylcysteine (Mucodyne), are reported. Fluidification of sputum with reduction in certain measurements of the viscosity of morning sputum aliquots, associated with improvement in the ability to cough up bronchial secretions, significant increase in sputum volume output, and improvement in ventilation (as estimated by the forced expiratory volume in one second), were observed in both trials as dose-related responses, with an increase in the ease of expectoration and a reduction in cough frequency and dyspnea. Therapy with S-carboxymethylcysteine was well tolerated, and there were no serious adverse effects, either immediate or delayed. We suggest that the effect of the drug in fluidifying sputum may be due to a mucoregulatory mechanism which reverses the sputum macromolecular disturbances seen in chronic bronchitis.

Check Tags: Female; Human; Male CONTROLLED TERM:

Administration, Oral

Adult

*Bronchitis: DT, drug therapy

Carbocysteine: AD, administration & dosage

Carbocysteine: PD, pharmacology *Carbocysteine: TU, therapeutic use

Chronic Disease Clinical Trials

*Cysteine: AA, analogs & derivatives

Forced Expiratory Volume

Humidity Middle Age

Respiratory Therapy *Sputum: DE, drug effects

Viscosity Vital Capacity

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

L48 ANSWER 25 OF 81 MEDLINE

75175470 MEDLINE ACCESSION NUMBER:

DOCUMENT NUMBER: 75175470 PubMed ID: 1134660

TITLE: [Studies of the clinical effectiveness of the mucolytic drug, S-carboxymethylcysteine, in the therapy of acute and

chronic bronchitis].

Indagine sull'efficacia clinica del mucolitico

S-carbossimetilcisteine nella terapia delle bronchiti acute

e croniche.

AUTHOR: Magliulo E; Bonizzoni D; Cattaneo E; Scevola D; Concia E

MINERVA MEDICA, (1975 Apr 4) 66 (25) 1187-97. SOURCE:

Journal code: 0400732. ISSN: 0026-4806.

Italy

PUB. COUNTRY:

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

Italian LANGUAGE:

Priority Journals FILE SEGMENT:

197508 ENTRY MONTH:

Entered STN: 19900310 ENTRY DATE:

Last Updated on STN: 19900310 Entered Medline: 19750820 Check Tags: Human; Male

CONTROLLED TERM: Acute Disease

Adult Aged

*Bronchitis: DT, drug therapy *Carbocysteine: TU, therapeutic use

Chronic Disease

*Cysteine: AA, analogs & derivatives *Expectorants: TU, therapeutic use

Middle Age

2387-59-9 (Carbocysteine); 52-90-4 (Cysteine) CAS REGISTRY NO .:

0 (Expectorants) CHEMICAL NAME:

MEDLINE L48 ANSWER 26 OF 81

MEDLINE 76154723 ACCESSION NUMBER:

PubMed ID: 769242 76154723 DOCUMENT NUMBER:

No demonstrable effect of S-carboxymethylcysteine on TITLE:

clearance of secretions from the human lung. Thomson M L; Pavia D; Jones C J; McQuiston T A

THORAX, (1975 Dec) 30 (6) 669-73. AUTHOR:

Journal code: 0417353. ISSN: 0040-6376. SOURCE:

ENGLAND: United Kingdom PUB. COUNTRY:

(CLINICAL TRIAL) DOCUMENT TYPE:

(CONTROLLED CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

197606 ENTRY MONTH:

Entered STN: 19900313 ENTRY DATE:

Last Updated on STN: 19980206 Entered Medline: 19760602

The mucolytic efficacy of S-carboxymethylcysteine has been assessed in a double-blind crossover trial in 16 patients with chronic obstructive bronchitis. No significant difference was found between drug and placebo after four or seven days' treatment in the rate of clearance of secretions from the lung. This was measured by external counting of previously inhaled polystyrene tracer particles tagged with technetium-99m (99mTc). Lateral scans across the right chest after inhaling the aerosol showed equal penetration of particles towards the periphery of the lung in drug and placebo runs; this indicated that the airways had not been cleared of mucus by the drug. There was no significant difference between drug and placebo runs in the number of coughs or the weight and radioactive content of sputum voided or raised at the end of the run by chest percussion and postural drainage. Ventilatory capacity was not significantly changed nor was there any subjective improvement in the patients as a result of taking the drug.

Check Tags: Human; Male CONTROLLED TERM:

Aged

*Bronchitis: DT, drug therapy Bronchitis: PP, physiopathology

Carbocysteine: AD, administration & dosage

*Carbocysteine: TU, therapeutic use

Clinical Trials

*Cysteine: AA, analogs & derivatives

Forced Expiratory Volume

Lung: AN, analysis
*Lung: SE, secretion

Middle Age

*Mucus: DE, drug effects Sputum: AN, analysis

Vital Capacity

CAS REGISTRY NO.: 2387-59-9 (Carbocysteine); 52-90-4 (Cysteine)

=> fil reg; s 2387-59-9
FILE 'REGISTRY' ENTERED AT 09:23:07 ON 09 JUL 2002
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STRUCTURE FILE UPDATES: 8 JUL 2002 HIGHEST RN 437701-77-4 DICTIONARY FILE UPDATES: 8 JUL 2002 HIGHEST RN 437701-77-4

TSCA INFORMATION NOW CURRENT THROUGH January 7, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

L49 2 2387-59-9 (2387-59-9/RN)

=> d ide 1-2

L49 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS

RN 25390-17-4 REGISTRY

CN Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Alanine, 3-[(carboxymethyl)thio]-, DL- (8CI)

CN DL-Cysteine, S-(carboxymethyl)-

OTHER NAMES:

CN 5-Amino-3-thiadihexanoic acid

CN DL-3-(Carboxymethylthio)alanine

S-(Carboxymethyl)-(RS)-cysteine

CN S-(Carboxymethyl)-DL-cysteine

CN S-(Carboxymethyl)cysteine

FS 3D CONCORD

DR **2387-59-9**

MF C5 H9 N O4 S

CI COM

CN

LC STN Files: ANABSTR, BEILSTEIN*, BIOSIS, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMLIST, CSCHEM, IPA, MEDLINE, NIOSHTIC, RTECS*, TOXCENTER, USAN, USPATFULL

(*File contains numerically searchable property data)

Other Sources: EINECS**

(**Enter CHEMLIST File for up-to-date regulatory information)

Structures you all Medline references

```
\begin{array}{c} ^{\rm NH_2} \\ | \\ _{\rm HO_2C-\,CH-\,CH_2-\,S-\,CH_2-\,CO_2H} \end{array}
```

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

```
20 REFERENCES IN FILE CA (1967 TO DATE)
20 REFERENCES IN FILE CAPLUS (1967 TO DATE)
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```
L49 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS
     638-23-3 REGISTRY
     L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Alanine, 3-[(carboxymethyl)thio]-, L- (6CI, 8CI)
OTHER NAMES:
     (L)-2-Amino-3-(carboxymethylthio)propionic acid
     (R)-S-(Carboxymethyl) cysteine
CN
     3-[(Carboxymethyl)thio]-L-alanine
CN
     Bronchokod
CN
     Carbocisteine
CN
     Carbocysteine
CN
     DF 1794Y
 CN
     L-(Carboxymethyl)cysteine
 CN
     LJ 206
 CN
 CN
     Muciclar
     Mucodyne
 CN
     Mucopront
 CN
     Rhinathiol
 CN
     Rhinatiol
 CN
      Rinatiol
 CN
      S-(Carboxymethyl)-(R)-cysteine
 CN
      S-(Carboxymethyl)-L-cysteine
 CN
      S-Carboxylmethyl-L-cysteine
 CN
      Thiodril
 CN
       2387-59-9
 AR
       STEREOSEARCH
  FS
       11139-64-3
  DR
       C5 H9 N O4 S
  MF
       STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
  CI
         BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN,
  LC
         CSCHEM, CSNB, DDFU, DRUGU, EMBASE, HODOC*, IFICDB, IFIPAT, IFIUDB,
         MRCK*, NIOSHTIC, PHAR, PHARMASEARCH, PROMT, RTECS*, TOXCENTER, ULIDAT,
         USPATFULL, VETU
           (*File contains numerically searchable property data)
       Other Sources: EINECS**, NDSL**, TSCA**, WHO
            (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry.

$$_{\mathrm{HO_{2}C}}$$
 s $_{\mathrm{NH_{2}}}^{\mathrm{R}}$ $_{\mathrm{CO_{2}H}}$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

523 REFERENCES IN FILE CA (1967 TO DATE)
23 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

523 REFERENCES IN FILE CAPLUS (1967 TO DATE) 13 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

=> fil medl hcapl

AUTHOR(S):

FILE 'MEDLINE' ENTERED AT 09:23:41 ON 09 JUL 2002

FILE 'HCAPLUS' ENTERED AT 09:23:41 ON 09 JUL 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

=> d ibib abs hitstr 148 27-81; fil hom

L48 ANSWER 27 OF 81 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1997:723411 HCAPLUS

DOCUMENT NUMBER: 128:43683

TITLE: Effects of fudosteine, a new mucoactive drug, on the increase in mucus secretion produced by secretagogues

> in human pulmonary mucoepidermoid carcinoma cells Kusano, K.; Nishiwaki, S.; Naito, H.; Takahashi, Y.; Tachibana, K.; Yokoyama, T.; Kai, H.; Miyata, T.

Central Research Laboratories, SS Pharmaceutical Co., CORPORATE SOURCE:

Ltd, Narita, 286, Japan

Pharmaceutical Sciences (1997), 3(8), 403-406 SOURCE:

CODEN: PHSCFB; ISSN: 1356-6881

Royal Pharmaceutical Society of Great Britain PUBLISHER:

DOCUMENT TYPE: Journal English LANGUAGE:

The purpose of this study was to clarify the effects of fudosteine ((-)-(R)-2-amino-3-(3-hydroxypropylthio) propionic acid), a new mucoactive drug, on mucus secretion in the human pulmonary mucoepidermoid carcinoma cell line NCI-H292. NCI-H292 cells produced hyaluronidase-resistant high-mol.-wt. glycoconjugates (HMWG), which were eluted in the void vol. on Superose 6HR column chromatog. ATP, bradykinin, and methacholine increased the basal secretion of [14C]glucosamine-labeled HMWG in NCI-H292 cells. Fudosteine significantly suppressed the HMWG secretion induced by ATP or bradykinin, but not that induced by methacholine. Other mucoactive drugs, such as ethylcysteine, ambroxol and carboxymethylcysteine, did not affect the increase in HMWG. These results indicate that fudosteine suppresses the mucin secretion induced by ATP and bradykinin in airway mucus-producing cancer cells.

IT 638-23-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(fudosteine effect on increase in mucus secretion produced by secretagogues in human pulmonary mucoepidermoid carcinoma cells, and comparison with other mucoactive drugs)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 28 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1997:12626 HCAPLUS

DOCUMENT NUMBER: 126:50995 TITLE:

Pharmaceutical composition containing acetylcysteine, carbocysteine or erdosteine in combination with a beta

2 agonist and an expectorant for the treatment of respiratory tract disorders

INVENTOR(S): PATENT ASSIGNEE(S): Holtshousen, Peter David Adcock Ingram Limited, S. Afr.; Ashworth, Stuart

David; Holtshousen, Peter David

SOURCE:

PCT Int. Appl., 16 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

English

LANGUAGE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT INTORUM		DATE
PATENT NO.	KIND DATE	APPLICATION NO. DATE
WO 9635452 W: AL, AM, ES, FI,	MD, MG, MK, MN,	JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
SG, SI RW: KE, LS, IE, IT, ZA 9603590 AU 9656556	MW, SD, SZ, UG, LU, MC, NL, PT, A 19961119 A1 19961129	AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, SE, BF, BJ, CF, CG, CI, CM, GA, GN 19960507 19960509 19950510 19950510 19960509
	, ugoful	in the treatment of respiratory tract

A pharmaceutical compn. useful in the treatment of respiratory tract disorders comprises as active ingredients; (a) acetylcysteine, AB carbocysteine, erdosteine or a pharmaceutically acceptable salt of any of these; and (b) a .beta.2-agonist, e.g. salbutamol, terbutaline; and (c) an expectorant, e.g. guaiphensin, sodium citrate, ammonium chloride. TΤ

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) 638-23-3, Carbocysteine (pharmaceutical contg. a cysteine deriv., .beta.2-agonist and an expectorant for treatment of respiratory tract disorders)

RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\mathrm{HO_{2}C}}$$
 S $_{\mathrm{NH_{2}}}^{\mathrm{R}}$ $_{\mathrm{CO_{2}H}}^{\mathrm{CO_{2}H}}$

L48 ANSWER 29 OF 81 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1997:154918 HCAPLUS

DOCUMENT NUMBER:

126:162255

TITLE:

Expectorant compositions

Hibi, Yoshiaki; Hirata, Takeo; Watanabe, Masazumi

INVENTOR(S): PATENT ASSIGNEE(S): Takeda Chemical Industries Ltd, Japan Jpn. Kokai Tokkyo Koho, 7 pp.

SOURCE:

CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. DATE KIND DATE PATENT NO. -----

JP 08337532 A2 19961224 JP 1995-147367 19950614

Expectorant compns. comprise mucus secretion-promoting herbal medicine and mucus viscosity adjusters/mucosa lubricants for the respiratory tract. An oral expectorant compn. comprises L-ethylcysteine-HCl 250, senega exts. 450, and aster exts. 450 mg with addn. of excipients.

IT 638-23-3, Carbocysteine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (expectorant compns.)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 30 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:309926 HCAPLUS

DOCUMENT NUMBER: 125:1027

TITLE: N-Acetyl-L-cysteine and its derivatives activate a Cl-

conductance in epithelial cells

AUTHOR(S): Koettgen, M.; Busch, A. E.; Hug, M. J.; Greger, R.;

Kunzelmann, K.

CORPORATE SOURCE: Physiol. Inst. Albert Ludwigs, Univ. Freiburg,

Freiburg, D-79104, Germany

SOURCE: Pfluegers Arch. (1996), 431(4), 549-555

CODEN: PFLABK; ISSN: 0031-6768

DOCUMENT TYPE: Journal LANGUAGE: English

N-Acetyl-L-cysteine (NAC) is a widely used mucolytic drug in patients with a variety of respiratory disorders including cystic fibrosis (CF). The beneficial effects of NAC are empirical and the exact mechanism of action in the airways remains obscure. In the present study the authors examd. the effects on whole-cell (wc) conductance (Gm) and voltage (Vm) of NAC and the congeners S-carboxymethyl-L-cysteine (CMC) and S-carbamyl-L-cysteine (CAC) and L-cysteine in normal and CF airway epithelial cells. L-Cysteine (1 mM) had no detectable effect. The increase of Gm (.DELTA.Gm) by the other compds. was concn. dependent and was (all substances at 1 mM) 3.8 (NAC), 4.2 (CMC) and 3.8 (CAC), resp. The changes in Gm were paralleled by an increased depolarization (.DELTA.Vm) when extracellular C1- concn. was reduced to 34 mM: under control conditions = 4.1 vs. 10.2 mV in the presence of NAC, CMC, CAC. the presence of NAC, CMC and CAC, the redn. in Cl-concn. was paralleled by a redn. of Gm by 2.1, indicating that all substances acted by increasing the C1- conductance. Anal. of intracellular pH did not reveal any changes by any of the compds. (1 mM). A Cl- conductance was also activated in HT29 colonic carcinoma and CF tracheal epithelial (CFDE) cells but not in CFPAC-1 cells, which do not express detectable levels of .DELTA.F508-CFTR, suggesting that the presence of CFTR may be a prerequisite for the redn. of Cl- currents. Next the authors examd. the ion currents in Xenopus oocytes microinjected with CFTR-cRNA. Water-injected oocytes did not respond to activation by forskolin and 3-isobutyl-1-methylxanthine (IBMX) (.DELTA.Gm = 0.08 .mu.S) and no current was activated when these oocytes were exposed to NAC or CMC. In contrast, in CFTR-cRNA-injected oocytes Gm was enhanced when intracellular cAMP (cAMP) was increased by forskolin and IBMX (Gm = 4.5 .mu.S). Gm was significantly increased by 0.74 .mu.S and 0.46 .mu.S when oocytes were exposed to NAC and CMC, resp. (both 1 mM). In conclusion, NAC and its

Page 26

congeners activate C1- conductances in normal and CF airway epithelial cells and hence induce electrolyte secretion which may be beneficial in CF patients. CFTR appears to be required for this response in an as yet unknown fashion.

638-23-3, S-Carboxymethyl-L-cysteine ΙT

RL: BAC (Biological activity or effector, except adverse); THU

(Therapeutic use); BIOL (Biological study); USES (Uses)

(N-Acetyl-L-cysteine and derivs. activate a Cl- conductance in normal and cystic fibrosis human airway epithelial cells in relation to CFTR)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\rm NH_2}^{\rm R.~CO_2H}$$

L48 ANSWER 31 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1996:625164 HCAPLUS ACCESSION NUMBER:

125:257189 DOCUMENT NUMBER:

Pharmaceutical composition containing a mucolytic agent and a bronchodilator for the TITLE: treatment of respiratory tract disorders

Treadwell, Cecil

Adcock Ingram Ltd., S. Afr. INVENTOR(S):

PATENT ASSIGNEE(S): S. African, 9 pp.

SOURCE: CODEN: SFXXAB

Patent DOCUMENT TYPE:

English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

INIBINI 2000			APPLICATION NO.	DATE
PATENT NO.	KIND	DATE	APPLICATION NO.	
		10050711	ZA 1994-155	19940111
ZA 9400155	Ω	19950711	77 1992-8567	19921106
PRIORITY APPLN. INFO	.: 	in unit	dosage form comprises	(a) a there

A pharmaceutical compn. in unit dosage form comprises (a) a therapeutic AB

dose of acetylcysteine (I) or carbocysteine or a pharmaceutically acceptable salt thereof; (b) a therapeutic dose of terbutaline (II) or a pharmaceutically acceptable salt thereof; and (c) one or more

pharmaceutically acceptable excipients. A capsule contained I 100-2000, II sulfate 1-5, diluent 5-200, glidants 0-15, and disintegrants 0-20 mg.

638-23-3, Carbocysteine

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compn. contg. mucolytic agent and

bronchodilator for treatment of respiratory tract disorders)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$_{
m HO_2C}$$
 S $_{
m NH_2}$

L48 ANSWER 32 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1996:35119 HCAPLUS

DOCUMENT NUMBER: 124:176944

TITLE: Preparation of S-(carboxymethyl)cysteine

Sato, Tadashi INVENTOR(S): Kojin Kk, Japan PATENT ASSIGNEE(S):

Jpn. Kokai Tokkyo Koho, 2 pp. SOURCE:

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

KIND DATE APPLICATION NO. DATE PATENT NO. JP 07267922 A2 19951017 JP 1994-82702 19940330 OTHER SOURCE(S): CASREACT 124:176944

The compd. (I), useful as an expectorant and an intermediates for biotins, is prepd. by treatment of L-cystine with ClCH2CO2H (II) in the presence of NaBH4. NaBH4 was gradually added to a mixt. of H2O, L-cystine, and II, which was previously adjusted to pH 8, under cooling and the reaction mixt. was further stirred at room temp. for 2 h to give 91% I.

638-23-3P, S-Carboxymethyl-L-cysteine ΙT

> RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of (carboxaymethyl) cysteine from cystine and C1CH2CO2H in presence of NaBH.beta.)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 33 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1995:997702 HCAPLUS

DOCUMENT NUMBER: 124:37727

Compound benproperine pharmaceutical compositions for TITLE:

respiratory infections

INVENTOR(S): Ye, Rongke

Baiyunshan Pharmaceutics Stock-Sharing Co., Ltd., PATENT ASSIGNEE(S):

Peop. Rep. China

Faming Zhuanli Shenqing Gongkai Shuomingshu, 4 pp. SOURCE:

CODEN: CNXXEV

DOCUMENT TYPE: Patent Chinese LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE 19950705 CN 1993-106616 A

AB Antiinflammatory, antitussive, and expectorant compns. for patients with respiratory infections comprise benproperine, carboxymethylcysteine and houttuynine at a ratio of 2:15:5. Capsules were formulated contg. benproperine 20, carboxymethyl cysteine 150, and houttuynine 50g.

IT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (compd. benproperine pharmaceutical compns. for respiratory infections)

638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) RN CN

Absolute stereochemistry.

HO₂C

L48 ANSWER 34 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1993:261047 HCAPLUS

ACCESSION NUMBER:

118:261047

Inhalation preparations containing carbocysteine DOCUMENT NUMBER: Kamijo, Shinji; Imai, Atsushi; Hibino, Kazuhide TITLE:

Kyorin Seiyaku Kk, Japan INVENTOR(S): PATENT ASSIGNEE(S): Jpn. Kokai Tokkyo Koho, 6 pp. SOURCE:

CODEN: JKXXAF

Patent DOCUMENT TYPE: Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

AB

APPLICATION NO. DATE KIND DATE PATENT NO. -----JP 1991-227099 19910906 _____ __________ A2 19930309 Aq. solns. (pH 6.0-7.5) contg. carbocysteine (I) and stabilizers are sealed with inert gas in a container for inhalation. I (5 g) was mixed with H2O, aq. NaOH, and 1 g Na citrate, adjusted to pH 7.0 with aq. NaOH, mixed with H2O to 1000 mL, charged into glass tube, the air purged with N, and sealed. The inhalation prepn. was stable at 50.degree. for .gtoreq.2 mo, vs. poor stability, without the citrate and N.

638-23-3, Carbocysteine TΤ

RL: BIOL (Biological study) (inhalation prepns. contg. stabilizers and, under inert gas)

638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) RNCN

Absolute stereochemistry.

L48 ANSWER 35 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1993:261046 HCAPLUS

ACCESSION NUMBER:

Inhalation preparations containing carbocysteine DOCUMENT NUMBER: TITLE:

Kamijo, Shinji; Imai, Atsushi; Hibino, Kazuhide

Kyorin Seiyaku Kk, Japan INVENTOR(S): PATENT ASSIGNEE(S): Jpn. Kokai Tokkyo Koho, 4 pp.

SOURCE: CODEN: JKXXAF

Patent DOCUMENT TYPE: Japanese LANGUAGE: 1

FAMILY ACC. NUM. COUNT:

Page 29

PATENT INFORMATION:

KIND DATE PATENT NO. APPLICATION NO. DATE ---------JP 05058887 A2 19930309 JP 1991-227098 19910906

Aq. carbocysteine (I) solns. (pH 6.0-7.5) are sealed with inert gas in a AB container for single-use inhalation. I (50 g) was mixed with H2O, adjusted to pH 7.0 with aq. NaOH, mixed with H2O to 1000 mL, charged into amples, the air purged with N, and sealed to make inhalation prepn. (contg. 2 mL/ample), which was stable at 50.degree. for .gtoreq.2 mo, vs. poor stability, when the soln. was charged (50 mL/each) in glass tube and sealed without N.

638-23-3, Carbocysteine ΙT

RL: BIOL (Biological study)

(inhalation prepns. contg., under inert gas, stable)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 36 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1994:8968 HCAPLUS

DOCUMENT NUMBER:

120:8968

TITLE:

Preparation of L-S-carboxymethylcysteine L-lysine salt

monohydrate as a drug

INVENTOR(S):

Argese, Maria; Bosone, Enrico; Clavenna, Gaetano;

Giani, Roberto

PATENT ASSIGNEE(S):

Dompe' Farmaceutici S.p.A., Italy

SOURCE:

Eur. Pat. Appl., 10 pp. CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.		DATE		APPLIC	CATION NO	•	DATE				
	EP 546272 EP 546272	A1			EP 199	92-117267	-	19921	1009			
	R: AT, BE, AT 134613 ES 2049700 JP 05310686	E T3	19960315 19960416		AT 199 ES 199	92-117267 92-117267		19921 19921	1009 1009	NL,	PT,	SE
PRIO	RITY APPLN. INFO					MI3338						
	L-S-carboxymethy				lt mond	ohydrate	(I)	was	prep	pd.	Thu	s,
	1 mol L-S-carbo contg. 1 mol of filtration and a mucolytic, brond (no data).	the la	tter follogf 5 vols.	wed b	y decol OH to q	lorizatio give 90%	n w I.	ith a I ha	activ as	ve C		
IT	638-23-3											
	RL: RCT (Reactar (salification	•	ith lysine	, in	prepn.	of drug)						
RN	638-23-3 HCAPL		-	-								
CN	L-Cysteine, S-(carboxy	methyl)- (9CI)	(CA II	NDEX NAME)					

Absolute stereochemistry.

$$_{\rm HO_2C}$$
 S $_{\rm NH_2}^{\rm R}$

L48 ANSWER 37 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1992:462714 HCAPLUS ACCESSION NUMBER:

117:62714

The influence of sulfhydryl-containing mucolytics on DOCUMENT NUMBER: the enzyme activity of human alveolar macrophages TITLE:

Mueller, L.; Wehle, K.; Kuepper, T.; Pfitzer, P. Abt. Zytopathol., Heinrich-Heine-Univ., Duesseldorf,

AUTHOR(S): CORPORATE SOURCE:

Atemwegs- Lungenkrankh. (1992), 18(3), 107-11 SOURCE:

CODEN: ATLUDF; ISSN: 0341-3055

Journal DOCUMENT TYPE:

Human sputa, bronchial secretions and lavages were stained cytochem. for LANGUAGE: several lysosomal enzymes after having been exposed to various concns. of N-acetylcysteine, carbocysteine, and mesna. A redn. of enzyme activity was obsd. in the samples treated with N-acetylcysteine in concn. >0.1%.

Concns. which can be achieved in bronchial mucus by systemic application did not reduce the enzyme activity.

638-23-3, Carbocysteine IT

RL: BIOL (Biological study)

(enzymes of lung lavage and sputum responses to, in human)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\rm HO_2C}$$
 s $_{\rm NH_2}^{\rm R}$ $_{\rm CO_2H}$

L48 ANSWER 38 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1992:113527 HCAPLUS

ACCESSION NUMBER:

116:113527 DOCUMENT NUMBER:

Pharmaceutical compositions containing TITLE:

phenylpropanolamine for a mucus secretagogue in the

upper airways

Phipps, Roger John INVENTOR(S):

Procter and Gamble Co., USA PATENT ASSIGNEE(S):

PCT Int. Appl., 33 pp. SOURCE:

CODEN: PIXXD2

Patent DOCUMENT TYPE: English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

NT INFORMATION:				рате
PATENT NO.	KIND DAT	E	APPLICATION NO	. DATE
WO 9117746	A1 199	11128	WO 1991-US3453	19910517
W: AU, CA, RW: AT, BE, AU 9179920		K, ES, FR,	GB, GR, IT, LU, AU 1991-79920	NL, SE 19910517

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EP 530311
                      Α1
                          19930310
                                         EP 1991-911164
                                                         19910517
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                          19931222 JP 1991-510443 19910517
                    Т2
    JP 05509300
                      Α
                          19920226
                                         ZA 1991-3831
                                                         19910521
    ZA 9103831
                      Α
                          19931109
                                         US 1992-893956
                                                         19920604
    US 5260073
                      A1
                           19950803
                                         AU 1995-20508
                                                         19950605
    AU 9520508
PRIORITY APPLN. INFO.:
                                      US 1990-526218
                                                         19900521
                                      WO 1991-US3453
                                                         19910517
```

AB Mucus secretion is induced in the upper airways of persons with sinusitis or otitis media (characterized by retention of thickened respiratory secretions) by administration of an effective amt. of d-(+)-norephedrine, l-(-)-norephedrine, or mixts. thereof. Oral formulations are presented, as are clin. effectiveness reports.

IT 638-23-3

RL: BIOL (Biological study)

(upper respiratory mucus secretion-inducing norephedrine compn. contg.)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 39 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1990:496164 HCAPLUS

DOCUMENT NUMBER: 113:96164

TITLE: Manufacture of S-(alkoxycarbonyl)methyl-L-cysteines

with tryptophan synthase

INVENTOR(S): Nakamura, Takeshi; Ishiwatari, Kenichi; Makiguchi,

Nobuyoshi

PATENT ASSIGNEE(S): Mitsui Toatsu Chemicals, Inc., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	JP 02072891	A2	19900313	JP 1988-221358	19880906
OTHE	R SOURCE(S):	MA	RPAT 113:96164		
AB	L-RO2CCH2SCH2CH(NH2)CC	2H (R = alkyl)	(I), useful as in	termediates for
				ne (II) and other	
					III) with HSCH2CO2R
					. Treatment of III
					mM with 0.9 units
				coli MT-10242 (FE	
	-		-		utoxycarbonyl)methyl-
					as treated similarly
	with thioglycoli	c acid	l instead of Bu	thioglycolate.	
ΙT	638-23-3P				
	RL: PREP (Prepar				

(prepn. of, intermediate for, alkoxycarbonylmethylcysteines as)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 40 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1991:150172 HCAPLUS

ACCESSION NUMBER:

DOCUMENT NUMBER:

TITLE:

Coating of drug particles with cellulose derivatives

Page 32

and acrylic polymers

INVENTOR(S):

Poli, Stefano; Moro, Luigi; Fiori, Achille; Natali,

Alberto

PATENT ASSIGNEE(S):

Poli Industria Chimica S.p.A., Italy

SOURCE:

Ger. Offen., 8 pp.

CODEN: GWXXBX

DOCUMENT TYPE: LANGUAGE:

Patent

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

LEMI INFORMATION	,		APPLICATION NO.	DATE
PATENT NO.	KIND	DATE		19891222
DE 3943242	711	19900628 19920612	FR 1989-17074	19891222 19881223
FR 2670112 RIORITY APPLN.	117		IT 1988-23075 . of a cellulose der	
XIONIII METERI	+	with a MIXT	. OI a CCITATO	The coa

Solid drugs are coated with a mixt. of a cellulose deriv. with a vinyl, acrylic and/or methacrylic and/or cyanoacrylic polymer. The coating is PRI for sustained-release, taste masking, stabilization, etc. particles of a AΒ mixt. contg. 2 kg dihydroerogocristine methanesulfonate and 13 kg inert excipient were coated with a mixt. of Aquacoat ACD-30 48, Eudragit NE30D (acrylic ester-methacrylic ester copolymer latex) 24, ionic surfactant 3, and water 25%.

638-23-3, Carbocysteine

RL: BIOL (Biological study)

(coating of particles of, with polymers)

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) RN

Absolute stereochemistry.

L48 ANSWER 41 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1991:35966 HCAPLUS

DOCUMENT NUMBER:

TITLE:

Temperature dependence of surface activity of films containing DPPC and selected drugs used in form of

AUTHOR(S): CORPORATE SOURCE: Pawelek, Janusz; Hanicka, Magdalena; Kurzawa, Ryszard

Inst. Catal. Surf. Chem., Pol. Acad. Sci., Krakow,

Journal

SOURCE:

Bull. Pol. Acad. Sci., Chem. (1990), Volume Date 1989,

37(9-12), 417-21

CODEN: BPACEQ; ISSN: 0239-7285

DOCUMENT TYPE:

Searched by Barb O'Bryen, STIC 308-4291

Page 33

LANGUAGE: English

During therapy based on inhalation, possible interactions between the applied inhalant and pulmonary surfactant should be taken into consideration. Therefore, it seemed useful to det. in vitro at different temps. to what extent such inhalants as Atrovent, Berotec, Bricanyl, Salbutamol, Mistabron and Ambroxol affect surface activity of the monolayer of dipalmitoyl lecithin (DPPC), the basic component of pulmonary surfactant. The measurements were performed in a Langmuir trough. The temp. varied from 20.degree. to 40.degree.. All the examd. inhalants were found to increase the surface activity of DPPC monolayer, esp. at 37.degree.. Only Mistabron decreased the surface activity of DPPC monolayer, which is the result of solubilization of DPPC mols.

ΙT 638-23-3

RL: BIOL (Biological study)

(aerosols contg., lung surfactant response to, temp.-dependent effects on dipalmitoylphosphatidylcholine membrane in evaluation of)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 42 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1991:57844 HCAPLUS

DOCUMENT NUMBER: 114:57844

Assay of aspartylglycosylaminase by high-performance TITLE:

liquid chromatography

Kaartinen, V.; Mononen, I. AUTHOR(S):

CORPORATE SOURCE: Dep. Clin. Chem., Kuopio Univ., Kuopio, SF-70210,

Finland

SOURCE: Anal. Biochem. (1990), 190(1), 98-101

CODEN: ANBCA2; ISSN: 0003-2697

DOCUMENT TYPE: Journal LANGUAGE: English

An aspartylglycosylaminase assay based on HPLC anal. of the substrate, aspartylglucosamine, and product, aspartate, is described. Aspartylglucosamine and aspartate are derivatized with

phenylisothiocyanate and resolved by reverse-phase HPLC. The detection limit for the compds. is 2 pmol. The method can be used for anal. of aspartylglycosylaminase activity in crude cell exts. and tissue samples.

IT 638-23-3

RL: ANT (Analyte); ANST (Analytical study)

(detn. of, HPLC method for)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$HO_2C$$
 S NH_2

L48 ANSWER 43 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1991:486955 HCAPLUS

ACCESSION NUMBER:

Page 34

Cook

DOCUMENT NUMBER:

115:86955

TITLE:

Morphologic changes of bronchial epithelial cells induced by sulfur dioxide during the recovery stage:

effect of the S-CMC and Ambroxol

AUTHOR(S):

Okamura, Takao; Satou, Shigeru; Aihara, Kaoru; Taga,

Fukutarou; Okamura, Kyuya

CORPORATE SOURCE:

Cent. Inst. Electron Microsc. Res., Nippon Med. Sch.,

SOURCE:

Nippon Kaimen Igakkai Zasshi (1990), 21(1-2), 19-35

CODEN: NKIZDR; ISSN: 0288-8262

DOCUMENT TYPE:

Journal Japanese

Continuous exposure of Wistar rat bronchial tree to 100 ppm SO2 revealed LANGUAGE: disintegration and loss of cilia and a decrease of mucin-secreting cells. These changes were most prominent after 3 days of exposure. After 1 wk, recovery of the damage was initiated and complete recovery was obsd. at 3 wk. The secretory granules prior to exposure to SO2 contained abundant PAS-pos. component presumably representing the neutral mucin. The mucin in the recovery stage consisted of basic mucin (Alcian Blue staining). The therapeutic effect of the S-CMC (carbocysteine) was represented by the prevention of degeneration. The cell protective effect of the Ambroxol was seen by the preservation of the cilia 3 days after exposure to SO2; however, no evidence for therapeutic effect of the agent was obsd. 1 wk after the exposure.

638-23-3, Carbocysteine · IT

RL: BIOL (Biological study)

(sulfur dioxide toxicity to bronchial epithelium prevention by)

638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) RN

Absolute stereochemistry.

$$_{\rm HO_2C}$$
 s $_{\rm NH_2}^{\rm R}$ $_{\rm CO_2H}$

L48 ANSWER 44 OF 81 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1990:125241 HCAPLUS

DOCUMENT NUMBER:

112:125241

TITLE:

Secretolytic agents for the prevention of

snoring

INVENTOR(S):

Reichert, Dietrich

PATENT ASSIGNEE(S):

SOURCE:

U.S., 5 pp. Division of U.S. Ser. No. 47,560.

CODEN: USXXAM

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE 19890320
US 4876283 DE 3317530 US 4831057 PRIORITY APPLN. IN	A A1 A	19891024 19841115 19890516	US 1989-325684 DE 1983-3317530 US 1987-47560 DE 1983-3317530 DE 1983-3317538 US 1984-609287 US 1987-47560 DE 1983-3317558	19830513 19870427 19830513 19830513 19840511 19870427 19830513

AB Antisnoring compns. for oral and local application in the nasal and pharyngeal cavities comprise a mucus-secreting agent (e.g. bromohexin) with diluents. Drops were manufd. from a mixt. contg. bromohexin 1.2 g, glycerol 1.0 mL, chlorobutanol 1.0 g, chamomile oil 0.2 g, and physiol. saline soln. to 100 g.

IT 638-23-3

RL: BIOL (Biological study)
 (snoring prevention by)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 45 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1990:77952 HCAPLUS

DOCUMENT NUMBER: 112:77952

TITLE: Preparation of S-carboxymethyl-L-cysteine

INVENTOR(S): Naijo, Shuichi; Inoue, Osami
PATENT ASSIGNEE(S): Showa Denko K. K., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 01193245	A2	19890803	JP 1988-17159	19880129
JP 2501852	B2	19960529	•	

L-Cystine is reduced by HSO3- to L-cysteine and S-sulfo-L-cysteine (I) at pH 5.5-6.0 and below the b.p. of the reaction system, the mixt. is directly treated with a monohaloacetic acid at pH 5-7.5 to give selectively S-carboxymethyl-L-cysteine (II), which is isolated; I in the mother liquor is hydrolyzed with a mineral acid to L-cystine for recycle. II is useful as an intermediate for pharmaceuticals (expectorants). Thus, a soln. of 2.00 g L-cystine and 2.69 g Na2SO3 in 300 g H2O was adjusted to pH 5.8 with concd. H2SO4 at 60.degree. and allowed to react to give a mixt. of 1.01 g L-cysteine and 1.68 g I; the mixt. was adjusted to pH 6.8 with 7.5N NaOH and treated with 1.21 g ClCH2CO2H at 60.degree. under N to give 1.34 g II. The mother liquor contg. I after sepn. of II was acidified with concd. H2SO4 and heated at 115.degree. to give 0.36 g L-cystine.

IT 638-23-3P, S-Carboxymethyl-L-cysteine

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of, use of bisulfite ions as reducing agents in, as

intermediate for pharmaceuticals)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl) - (9CI) (CA INDEX NAME)

HO2C

L48 ANSWER 46 OF 81 HCAPLUS COPYRIGHT 2002 ACS

Cook

1990:93732 HCAPLUS ACCESSION NUMBER:

112:93732

Oral N-acetylcysteine or S-carboxymethylcysteine DOCUMENT NUMBER: inhibit cigarette smoke-induced hypersecretion of TITLE:

mucus in rat larynx and trachea in situ

Rogers, D. F.; Turner, N. C.; Marriott, C.; Jeffery, AUTHOR(S):

Natl. Heart Lung Inst., Brompton Hosp., London, SW3 CORPORATE SOURCE:

6HP, UK

Eur. Respir. J. (1989), 2(10), 955-60 SOURCE:

CODEN: ERJOEI; ISSN: 0903-1936

Journal DOCUMENT TYPE: English

Two weeks exposure of rats to cigarette smoke (CS) significantly increased LANGUAGE: the secretion of fucose-contg. glycoconjugates above normal in an in situ prepn. of larynx and trachea. After equilibration mean basal secretion in CS-exposed rats was 24 .mu.g (per 30 min collection) which was 8 times higher than that of unexposed animals. N-Acetylcysteine (NAC) or S-carboxymethylcysteine (SCMC) given as 1% of the drinking water, before and after daily exposure to CS, significantly inhibited the development of the CS-induced increase in fucose secretion reducing the mean for basal secretion in each group to 7 and 5 .mu.g, resp. Neither NAC nor SCMC had significant effects on baseline glycoconjugate secretion in control animals. Albumin was inconsistently present in the secretions of both control and CS-exposed animals, whereas in those exposed to CS and also given one of the two cysteine derivs. there was a consistent increase in albumin transudation.

638-23-3 IT

RL: BIOL (Biological study)

(cigarette smoke effect on mucus secretion by larynx and trachea response to)

638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) RN CN

Absolute stereochemistry.

L48 ANSWER 47 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1988:556294 HCAPLUS ACCESSION NUMBER:

109:156294 DOCUMENT NUMBER:

Expectorants containing carbocysteine and TITLE:

syrups. Kamijo, Shinji; Imai, Atsushi; Hibino, Kazuhide INVENTOR(S):

Kyorin Pharmaceutical Co., Ltd., Japan PATENT ASSIGNEE(S):

Jpn. Kokai Tokkyo Koho, 4 pp.

SOURCE: CODEN: JKXXAF

Patent DOCUMENT TYPE: Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 63156719	A2	19880629	JP 1986-303462	19861219
JP 08013737	B4	19960214		

AB Expectorant syrups are prepd. contg. carbocysteine (I) and sugar alcs., which stabilize I. An aq. I soln. (5 % by wt./vol.) was mixed with 40 % by wt./vol. D-sorbitol (II) and left at 50.degree. for 30 days to show pH change -0.03, coloration (measured by colorimeter) 2.4, and residual I 97%, vs. -0.94, >50, and 42%, resp., for a control contg. glucose instead of I. A syrup was prepd. consisting of I 50, sorbic acid 1.0, II 400, caramel 0.6 g, aq. NaOH, fruit essence, and H2O.

TΤ 638-23-3

RL: BIOL (Biological study)

(expectorant contq. sugar alcs. as sweeteners and, stability in relation to)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl) - (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 48 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1988:89201 HCAPLUS

DOCUMENT NUMBER: 108:89201

Morphologic changes of bronchial epithelium induced by TITLE:

sulfur dioxide and protective effect of the S-CMC

administration

Takao, Okamura; Shigeru, Satou; Kaoru, Aihara; AUTHOR(S):

Kouichi, Takagi; Kyuya, Okamura

Cent. Inst. Electron Microsc. Res., Nippon Med. Sch., CORPORATE SOURCE:

Japan

Nippon Kaimen Igakkai Zasshi (1987), 18(1-2), 96-112 SOURCE:

CODEN: NKIZDR

DOCUMENT TYPE: Journal LANGUAGE: Japanese

In rats exposed to 100 ppm SO2 for 6, 24, or 48 h, the no. of ciliated bronchial epithelial cells decreased below controls; the cells desquamated after 2-3 days. The no. of goblet cells in the bronchi increased after 6 h and desquamated after 1-3 days. After 7-day recovery, some regeneration of the damage was obsd. Oral pretreatment of the rats with 200 mg carbocysteine (S-CMC)/day for 5 days and during the SO2 exposure prevented most of the damage. Thus, carbocysteine protects the bronchial epithelium from SO2 damage.

638-23-3, Carbocysteine TΨ

RL: BIOL (Biological study)

(bronchi epithelium protection by, against sulfur dioxide inhalation toxicity)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

$$_{
m NH}_{
m 2}$$

L48 ANSWER 49 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1987:432958 HCAPLUS

DOCUMENT NUMBER:

107:32958

Cook

TITLE:

Comparison of the course of regeneration of changes induced in the respiratory epithelium by the oral

administration of two different mucolytics

Konradova, V.; Vavrova, V.; Sulova, J.

AUTHOR(S): CORPORATE SOURCE:

SOURCE:

Karlova Univ., Prague, Czech. Stud. Pneumol. Phtiseol. Cech. (1987), 47(1-2), 86-100

CODEN: SPPCAC; ISSN: 0371-2222

DOCUMENT TYPE:

Journal

The authors investigated the course of regeneration after changes induced LANGUAGE: in the rabbit tracheal epithelium by oral administration of 1 dose of 2 mucolytic prepns., Bromhexine and Mucopront. Both mucolytics cause marked damage of the goblet cells of the respiratory epithelium. After rapid evacuation of the secretion, they degenerate. Bromhexine stimulates the massive differentiation of new mucus-secreting elements and formation of intraepithelial mucous glands. Changes induced in the tracheal epithelium by the action of the mucolytics do not recede within 3 days. Regeneration of goblet cells occurs sooner after administration of Bromhexine than after Mucopront. After Bromhexin there is also a more rapid decrease in the no. of stimulated and degenerated goblet cells in the epithelium. Seventy-two hours after administration of the 2 prepns., the no. of mucus-filled and degenerated goblet cells in the 2 exptl. groups does not differ. In both instances, in the epithelium 10% degenerated goblet cells remain. Administration of the 2 substances causes local disorders of mucus flow in the respiratory pathways. The impaired self-cleaning capacity of the epithelium is restored after administration of Mucopront in 3 days, after Bromhexine bacteria and condensed layers of mucus from the area of the ciliary border do not disappear within 72 h. The persisting disorder of the self-cleaning capacity of the epithelium is obviously due to the subsequent hyperplasia of the goblet cells and the appearance of the intraepithelial mucous glands.

638-23-3 ΙT

RL: BIOL (Biological study)

(respiratory epithelium regeneration after administration of)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 50 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1986:618683 HCAPLUS

ACCESSION NUMBER: DOCUMENT NUMBER:

TITLE:

105:218683 Preclinical and clinical investigation on combination effects of expectorants in chemotherapy of infectious respiratory diseases

AUTHOR(S):

Imaoka, Makoto

CORPORATE SOURCE:

Dep. Int. Med., Shimane Prefect. Cent. Hosp., Izumo,

693, Japan

SOURCE:

Chemotherapy (Tokyo) (1986), 34(3), 262-70

CODEN: NKRZAZ; ISSN: 0369-4682

DOCUMENT TYPE:

Journal LANGUAGE: Japanese

Mice were orally treated with rifampicin (I) [13292-46-1], ampicillin [69-53-4], orcephalexin [15686-71-2] alone or in combination with expectorants ambroxol (II) [18683-91-5], carbocysteine [638-23-3], or serratiopeptidase [37312-62-2]. After combination treatment with expectorants peak blood levels of the antibiotics increased in serum, lung, liver, and kidney. After combination of I plus II, the antibiotic concns. increased in serum and lung; the peak level increased by 46-137%. The results are discussed in terms of chemotherapy of infectious respiratory disease.

TΤ 638-23-3

RL: BIOL (Biological study)

(respiratory tract infection therapy with antibiotics and)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 51 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1986:218907 HCAPLUS

DOCUMENT NUMBER:

104:218907

TITLE:

Regeneration of changes induced in rabbit tracheal epithelium by a single oral dose of a mucolytic agent

Konradova, V.; Vavrova, V.; Sulova, J.

AUTHOR(S): CORPORATE SOURCE:

Fac. Pediatr., Charles Univ., Prague, Czech.

SOURCE:

Folia Morphol. (Prague) (1986), 34(1), 52-8

CODEN: FMORAO; ISSN: 0015-5640

DOCUMENT TYPE:

Journal

LANGUAGE:

English

In rabbits, carbocysteine [638-23-3] (100 mg, orally) did not have much effect on the ciliated cells, but it markedly stimulated and damaged the goblet cells. The percentage of degenerated goblet cells peaked in 60 min. Seventy-two h after administration of the mucolytic agent 26% of the stimulated and 10% of degenerated goblet cells were left in the epithelium. The excessive amt. of secretory material released into the airway lumen caused local disturbances of the mucus flow which were still pronounced 24 h after the drug administration and had not disappeared completely by the end of the exptl. period (72 h).

IT638-23-3

RL: BIOL (Biological study)

(toxiciity of, to trachea epithelium)

ŔŊ 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Cook

HO₂C NH₂

L48 ANSWER 52 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1986:88964 HCAPLUS ACCESSION NUMBER:

104:88964 DOCUMENT NUMBER:

Highly-pure S-(carboxymethyl)-L-cysteine TITLE:

hydrocholoride

Ajinomoto Co., Inc., Japan PATENT ASSIGNEE(S): Jpn. Kokai Tokkyo Koho, 2 pp. SOURCE:

CODEN: JKXXAF

Patent DOCUMENT TYPE: Japanese

LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE PATENT NO. KIND DATE _____ JP 1983-180030 19830928 JP 60072857 A2 19850424

Title compd., the hydrochloride salt of an expectorant (no data), was prepd. in high purity by crystg. S-(carboxymethyl)-L-cysteine AB (I) as the HCl salt from its soln. contg. L-cystine (II). Thus, a soln. of 200 g I contg. 1% II in aq. HCl was heated at 70.degree., seeded with I.HCl, cooled, filtered, and the filtrate neutralized with NaOH to give 120 g I of 99.5% purity (II content .ltoreq. 0.20%).

638-23-3P ΙT

RL: PUR (Purification or recovery); PREP (Preparation) (purifn. of)

638-23-3 HCAPLUS

RNL-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

R CO2H HO₂C NH2

L48 ANSWER 53 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1986:88962 HCAPLUS ACCESSION NUMBER:

104:88962 DOCUMENT NUMBER:

Purification of S-(carboxymethyl)-L-cysteine TITLE:

Ajinomoto Co., Inc., Japan PATENT ASSIGNEE(S): Jpn. Kokai Tokkyo Koho, 2 pp. SOURCE:

CODEN: JKXXAF

Patent DOCUMENT TYPE: Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

AΒ

APPLICATION NO. DATE PATENT NO. KIND DATE -----A2 19850424 JP 1983-180028 19830928 JP 60072855 The title compd. (I), useful as an expectorant (no data), was purified by seeding solns. contg. both I and cystine (II) with II and sepg. II by adjusting to a pH close to its isoelec. point. Thus, a soln. of 40 g I in aq. NaOH contg. 2 wt.% of II was adjusted to pH 6.5, 0.2 g II added, and the resulting mixt. heated at 50.degree. for 4 days to give 38 g I of .gtoreq.99.5% purity.

IT 638-23-3P

RL: PUR (Purification or recovery); PREP (Preparation)
 (purifn. of, as expectorant)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 54 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1986:51102 HCAPLUS

DOCUMENT NUMBER: 104:51102

TITLE: S-carboxymethyl-D-cysteine

PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan SOURCE: Jpn. Kokai Tokkyo Koho, 2 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 60069063	A2	19850419	JP 1983-178757	19830927
TP 03040022	R4	19910617		

AB Title compd. (I), useful as an expectorant (no data), was purified by adsorbing L-cystine (II) with strongly acidic cation exchangers from the acidic soln. of I contg. II. Thus, HCl soln. of I (pH = 0.2) contg. II was eluted through strongly acidic cation exchanger (SKIB, H type) and the eluant neutralized to give I in 99.5% purity.

IT 638-23-3P

RL: PUR (Purification or recovery); PREP (Preparation)
 (purifn. of, as expectorant)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 55 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1987:428377 HCAPLUS

DOCUMENT NUMBER: 107:28377

TITLE: Process for the preparation of zinc carbocysteinate

INVENTOR(S): Buxade Vinas, Antonio

PATENT ASSIGNEE(S): Laboratorios Vinas S. A., Spain

SOURCE: Span., 7 pp.

CODEN: SPXXAD

DOCUMENT TYPE: Patent

Cook

Spanish LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ES 524820	A1	19850201	ES 1983-524820	19830808
GB 2146324	A1	19850417	GB 1984-4634	19840222
GB 2146324	B2	19870218	FR 1984-12271	19840727
FR 2550446	A1	19850215	FR 1964-12271	15010.
FR 2550446	B1	19871211	JP 1984-161492	19840731
JP 60054354	A2	19850328 19861021	US 1984-636201	19840731
US 4618625	A A1	19850228	DE 1984-3428399	19840801
DE 3428399		19030220	ES 1983-524820	19830808
ORITY APPLN. INFO	• •		- 3	gent (no c

Zn carbocysteinate (1:2) (I), useful as a mucoserous agent (no data), is PRIORITY APPLN. INFO.: prepd. by salification of carbocysteine (II) with ZnO, Zn(OH)2, or another In salt in approx. stoichiometric proportions in one or more polar solvents. An aq. soln. of 44 g Zn (OAc)2 was added to an aq. suspension of 35.8 g II, and the mixt. was refluxed for 2 h to ppt. 35 g I. 638-23-3, Carbocysteine RL: BIOL (Biological study)

(salt exchange of, with zinc salts) 638-23-3 HCAPLUS

RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$_{
m R}$$
 $_{
m CO_2H}$ $_{
m NH_2}$

L48 ANSWER 56 OF 81 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1985:523924 HCAPLUS

DOCUMENT NUMBER:

103:123924

TITLE:

Cysteine derivatives

INVENTOR(S):

Puricelli, Laura

PATENT ASSIGNEE(S):

Magis Farmaceutici S.r.l., Italy

SOURCE:

Eur. Pat. Appl., 18 pp.

CODEN: EPXXDW Patent

DOCUMENT TYPE:

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 143399 EP 143399 EP 143399 R: AT, BE, AT 35536 US 4559360 PRIORITY APPLN. INFO	E A	19850605 19860723 19880706 , FR, GB, L 19880715 19851217	EP 1984-113796 I, LU, NL, SE AT 1984-113796 US 1984-673619 IT 1983-23844 EP 1984-113796	19841115 19841115 19841121 19831123 19841115

Ι

AB Cysteine derivs. RO2CCH2SCH2CH(NHR1)CO2R2 (R, R1, R2 = H, acetylsalicyloyl) were prepd. as mucolytics, antipyretics, analgesics, and inflammation inhibitors. Thus, L-HO2CCH2SCH2CH(NH2)CO2H was N-acylated with 2-AcOC6H4COCl in THF to give cysteine deriv. I. The toxicity of I in rats was LD50 4,800 mg/kg (oral). Antiinflammatory, antipyretic, and antibronchial activities of the above cysteine derivs. are discussed.

IT 638-23-3

RL: RCT (Reactant)

(benzoylation of, with acetoxybenzoyl chloride)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 57 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1985:553780 HCAPLUS

DOCUMENT NUMBER: 103:153780

TITLE: Comparison of the effect of three oral mucolytics on

the ultrastructure of the tracheal epithelium in

rabbits

AUTHOR(S): Konradova, Vaclava; Vavrova, V.; Sulova, J.

CORPORATE SOURCE: Lab. Electron Microsc., Res. Inst. Child Dev., Prague,

150 00, Czech.

SOURCE: Respiration (1985), 48(1), 50-7

CODEN: RESPBD; ISSN: 0025-7931

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The effect of the application of a single oral dose of 100 mg of three mucolytic agents [N-acetylcysteine [616-91-1], carbocysteine [638-23-3] and 2-mercaptoethane sulfonic acid [3375-50-6]] on the ultrastructure of the tracheal epithelium of healthy rabbits was studied.

Due to the function of oral mucolytics, the goblet cells were overstimulated and injured. 2-Mercaptoethane sulfonic acid was the least irritating substance. Acetylcysteine produced the most pronounced injury of the globlet cells, followed by rapid differentiation of these cells. Oral administration of all mucolytics studied caused local impairment of

mucus flow in the airways.

IT 638-23-3

RL: BIOL (Biological study)

(trachea epithelium toxicity from)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

CO2H s · HO₂C NH2

L48 ANSWER 58 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1985:84405 HCAPLUS ACCESSION NUMBER:

102:84405 DOCUMENT NUMBER:

Oral antisnoring agent TITLE: Reichert, Dietrich INVENTOR(S):

PATENT ASSIGNEE(S): Spain

Ger. Offen., 17 pp. SOURCE:

CODEN: GWXXBX

Patent DOCUMENT TYPE: German LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
		10041115	DE 1983-3317558	19830513
DE 3317558	A1	19841115		19840509
AU 8427807	A1	19841115	AU 1984-27807	19840309
AU 574360	B2	19880707		
EP 125634	À1	19841121	EP 1984-105287	19840510
EP 125634	B1	19900926		
		. FR, GB, IT	, LI, NL, SE	
R: AT, BE,	,	19901015	AT 1984-105287	19840510
AT 56879	E		JP 1984-96305	19840514
JP 60034910	A2	19850222		19870427
US 4831057	Α	19890516	US 1987-47560	
			DE 1983-3317530	19830513
PRIORITY APPLN. INFO	• •		DE 1983-3317558	19830513
			EP 1984-105287	19840510
			US 1984-609287	19840511
				1

An oral antisnoring agent contains compds. with secretolytic AB effects or effects on mucous-membrane secretion prodn., carriers and diluents. The active compd. regulate and normalize mucus viscosity, decrease mucus adhesion by activating surfactant properties of the secretion, stimulation of mucus prodn., and activate the mucociliary function. Capsules were prepd. from a mixt. of ambroxol-HCl [23828-92-4] 10.0, cornstarch 24.0, and Aerosil 200 0.4 g; each capsule contained 170 mg mixt.

ΙT 638-23-3

RL: BIOL (Biological study)

(oral antismoring pharmaceuticals contg.)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\mathrm{R}}^{\mathrm{CO}_{2}\mathrm{H}}$$

L48 ANSWER 59 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1985:84406 HCAPLUS ACCESSION NUMBER:

102:84406 DOCUMENT NUMBER:

TITLE:

Antisnoring agent Reichert, Dietrich INVENTOR(S):

PATENT ASSIGNEE(S):

Spain

SOURCE:

Ger. Offen., 17 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA'	TENT NO.		KIND	DATE	APPLICATION NO.	DATE
	3317530 1216232		A1 A1	19841115 19870106	DE 1983-3317530 CA 1983-429349	19830513 19830531
AU	8427807 574360		A1 B2	19841115 19880707	AU 1984-27807	19840509
ZA	8403502		A	19841224	ZA 1984-3502	19840509
EP	1223210 125634		A1 A1	19870623 19841121	CA 1984-453961 EP 1984-105287	19840509 19840510
EP	125634 R: AT,	BE, C	B1 H, DE	19900926 , FR, GB,	IT, LI, NL, SE	
	56879 60034909)	E A2	19901015 19850222	AT 1984-105287 JP 1984-96304	19840510 19840514
	662734 4831057		A A	19871030 19890516	CH 1984-4619	19840926 19870427
US	4876283	TNEO	A	19891024	US 1989-325684	19890320
PRIORIT	Y APPLN.	INFO.:			DE 1983-3317530 DE 1983-3317538	19830513 19830513
		•			DE 1983-3317558 EP 1984-105287	19830513 19840510
					US 1984-609287 US 1987-47560	19840511 19870427
			_			

AB An antisnoring agent for topical application to the nose or throat contains a secretolytic together with mucous membrane-compatible carriers or diluents. The compns. are in spray or inhalation soln. forms. The active secretolytics regulate and normalize mucus viscosity, decrease mucus adhesion by activation of surfactant properties, stimulate serous mucus prodn., and activate the mucociliary function. A nasal compn. was prepd. from a mixt. of ambroxol-HCl [23828-92-4] 10.0, glycerol [56-81-5] 1.0 mL, benzalkonium chloride 1.0 g and physiol. saline soln. to 100 mL for 1/2 - 1 mL doses.

IT 638-23-3

RL: BIOL (Biological study)

(nasal antisnoring pharmaceuticals contg.)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 60 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1985:25034 HCAPLUS

DOCUMENT NUMBER:

102:25034

TITLE:

S-Carboxymethylcysteine derivatives with therapeutic activity and pharmaceutical compositions containing

them

PATENT ASSIGNEE(S):

Istituto Biochimico Pavese S.p.A., Italy

SOURCE:

Belg., 17 pp. CODEN: BEXXAL

DOCUMENT TYPE:

Patent French

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
BE 899777 ES 532870 CH 661511 FR 2549057 FR 2549057	A1 A1 A A A1 B1	19840917 19850616 19870731 19850118 19870116	BE 1984-213024 ES 1984-532870 CH 1984-2628 FR 1984-8519	19840529 19840526 19840529 19840530
PRIORITY APPLN. INFO.	:		IT 1983-21358	19830330

S-Carboxymethylcysteine derivs. I (R = H, alkali or alk. earth metal, or org. or inorg. base or basic amino acid or antibiotic residue; R1 = H, AB alkali or alk. earth metal) were prepd. Thus, an aq. soln. of S-carboxymethylcysteine was neutralized with NaOH, 2-thiophenecarbonyl chloride in AcOEt added at 0-5.degree., and the mixt. kept at 2 h at 30-40.degree. to give I (R = R1 = $^{\circ}$ H) (II). The Ca salt of II had higher mucolytic activity than acetylcysteine in mice.

IT

RL: RCT (Reactant)

(acylation of, with thiophenecarbonyl chloride)

638-23-3 HCAPLUS

RN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

638-23-3DP, derivs. IT

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 61 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1983:482288 HCAPLUS

DOCUMENT NUMBER:

TITLE:

Change in glycoprotein composition in tracheal

Cook 09/868106

Page 47

submucosal glands by S-carboxymethylcysteine treatment

Takeda, Hiroshi; Ohkura, Yasufumi; Misawa, Miwa; AUTHOR(S):

Yanaura, Saizo

Sch. Pharm., Hoshi Univ., Tokyo, 142, Japan CORPORATE SOURCE:

SOURCE: Nippon Yakurigaku Zasshi (1983), 82(1), 19-25

CODEN: NYKZAU; ISSN: 0015-5691

DOCUMENT TYPE: Journal Japanese LANGUAGE:

The mechanism for the expectorant effect of S-carboxymethylcysteine (I) [638-23-3] was studied histol., and histochem. using isolated canine trachea. Following I treatment, the no. of total glycoprotein-contg. goblet cells (GC) did not change. The nos. of acid

glycoprotein (II)-,. neutral glycoprotein (III)-, and sulfated glycoprotein (IV)-contg. GC were also unaltered in a concn. range of 10-7 to 10-4M. On the other hand, the ratio of the acinar inner diam. of submucosal glands (SG) to the tracheal wall thickness was increased with 10-5 and 10-4M of I and the thickness of the acini of SG decreased with 10-4M I. Although the ratio of the nos. of II- to III-contg. glandular cells and II content in the cells did not change, the no. of IV-contg.

glandular cells significantly decreased concn.-dependently. Apparently, I

had a selective secretagogic action on SG, and an action which alters the compn. of II, a chief viscous factor, in the mucous granules of SG.

638-23-3 IT

> RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)

(glycoproteins of tracheal submucosal glands response to)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 62 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1983:149605 HCAPLUS

98:149605 DOCUMENT NUMBER:

TITLE: Pharmaceutical composition containing

S-carboxymethylcysteine and sobrerol

INVENTOR(S): Massaroli, Giangiacomo

PATENT ASSIGNEE(S): Poli Industria Chimica S.p.A., Italy

SOURCE: Fr. Demande, 9 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent

LANGUAGE: French FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE FR 2506614 A1 19821203 FR 1982-9236 19820527 FR 2506614 В1 19860314 DE 1982-3219994 19820527 DE 3219994 A1 19830127 PRIORITY APPLN. INFO.: IT 1981-22000 19810528 GI

sobrerol (I) [498-71-5] is combined with the mucolytic agent AB S-carboxymethylcysteine (II) [638-23-3] to improve its antiinflammatory-mucolytic activity, to reduce the viscosity of bronchial secretions, and to favor the regeneration of bronchial mucus. A syrup was prepd. contg. II 5, I 0.8, sucrose 42, Me p-hydroxybenzoate 0.15, NaH2PO4 0.5, NaOH 1.11, flavor 0.0665, and H2O to 100 g. Capsules and suppositories and other dosage forms contg. I and II were also prepd.

638-23-3 TT

RL: BIOL (Biological study)

(antitussive compns. contg. sobrerol and)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 63 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

. 1982:608359 HCAPLUS

DOCUMENT NUMBER:

97:208359

TITLE:

Effect of oral administration of the mucolytic agent Mucopront on the ultrastructure of the epithelium of

the respiratory passages

AUTHOR(S):

Konradova, V.; Vavrova, V.; Sulova, J. Fak. Detskeho Lek., KU, Prague, Czech.

CORPORATE SOURCE: SOURCE:

Cesk. Pediatr. (1982), 37(9), 497-500, 2 plates

CODEN: CEPEA3; ISSN: 0069-2328

DOCUMENT TYPE:

Journal

LANGUAGE:

Czech

At 20 and 60 min after a single oral administration of 100 mg Mucopront (carbocysteine) [638-23-3] to rabbits, no effect was obsd. on the ultrastructure of the ciliated cells of the tracheal epithelium. Goblet cells, however, were markedly affected. The excessive amt. of mucus released from the goblet cells impaired the self-cleaning ability of the tracheal epithelium.

638-23-3 IT

RL: BIOL (Biological study)

(respiratory tract epithelium ultrastructure response to)

638-23-3 HCAPLUS

RN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

$$R = CO_2H$$
 NH_2

L48 ANSWER 64 OF 81 HCAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 1981:575817 HCAPLUS

DOCUMENT NUMBER: 95:175817

TITLE: Pharmaceutical compositions with mucolytic,

bronchosecretolytic and antibronchospastic activity

PATENT ASSIGNEE(S): Dompe Farmaceutici S.p.A., Italy

SOURCE: Belg., 6 pp. CODEN: BEXXAL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

האתראות אוה V

PATENT NO. KIND DATE APPLICATION NO. DATE

BE 887922 Al 19810701 BE 1981-204105 19810312

PRIORITY APPLN. INFO.: IT 1980-20699 19800317

AB (.+-.)-Lysine (-)-carboxymethylcysteinate (I) [79458-68-7], prepd. by treating (-)-S-carboxymethylcysteine [638-23-3] with (.+-.)-lysine-HCl [70-53-1], has a high mucolytic, bronchosecretolytic, and antibronchospastic activity and high bioavailability, as compared to carboxymethylcysteine. I can be used in the form of tablets, capsules, syrups, or parenteral solns.

IT 638-23-3

RL: RCT (Reactant)

(reaction of, with lysine hydrochloride)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 65 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1982:173826 HCAPLUS

DOCUMENT NUMBER: 96:173826

TITLE: Tissue distribution of S-carboxymethylcysteine in the

rat: concentration in mucus-producing organs including

the prostate

AUTHOR(S): Bodmer, Judith L.; Waring, Rosemary H.

CORPORATE SOURCE: Dep. Biochem., Univ. Birmingham, Birmingham, B15 2TT,

UK

SOURCE: Biochem. Soc. Trans. (1981), 9(6), 549-50

CODEN: BCSTB5; ISSN: 0300-5127

DOCUMENT TYPE: Journal LANGUAGE: English

AB Autoradiog. examn. of the distribution of tissue labeling in rats given 14C-labeled S-carboxymethylcysteine (I) [638-23-3] (100 mg/kg, orally) indicated that I or a metabolite has some affinity for bronchopulmonary tissue, but that concns. in other mucus-producing organs may be as high if not higher than those in the lung. This may have therapeutic implications, particularly in the treatment of chronic

prostatitis, where the mucolytic action of I might be effective.

IT 638-23-3

RL: BPR (Biological process); BIOL (Biological study); PROC (Process)

(pharmacokinetics of, in mucus-producing tissues)

RN 638-23-3 HCAPLUS.

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\rm NH_2}^{\rm R}$$

L48 ANSWER 66 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1980:472302 HCAPLUS ACCESSION NUMBER:

93:72302 DOCUMENT NUMBER:

Cysteine derivatives TITLE:

Dong, Le Hao; Coquelet, Claude INVENTOR(S):

Laboratoires Chauvin-Blache S. A., Fr. PATENT ASSIGNEE(S):

Fr. Demande, 12 pp. Addn. to Fr. Demande 2,266,502. SOURCE:

CODEN: FRXXBL

Patent DOCUMENT TYPE: French LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE KIND DATE PATENT NO. _____ 19780712 FR 1978-20878 19800208 FR 2430945 Α2

GΙ

Cysteines I (R = H, C1-8 alkyl; R1 = .alpha.-thenoyl, CH2CO2H) were prepd. as bronchial protectors. Thus, N-.alpha.-thenoyl-L-cysteine Me ester was AB S-acylated with .alpha.-thenoyl chloride to give 70% cysteine II. I (R = H, R1 = CH2CO2H) (III) was also prepd. Data are given on the treatment of rats exposed to SO2 gas with II (360 mg/kg) and III (400 mg/kg).

638-23-3 ΙT

RL: RCT (Reactant)

(N-acylation of, with thenoyl chloride)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 67 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1981:57721 HCAPLUS ACCESSION NUMBER:

94:57721 DOCUMENT NUMBER:

TITLE:

The effect of mucolytic agents on the rheologic and

transport properties of canine tracheal mucus

AUTHOR(S):

Puchelle, E.; Sadoul, P.

CORPORATE SOURCE:

Unite Rech. Physiopathol., INSERM, Vandoeuvre-les-Nancy, 54500, Fr.

SOURCE:

Am. Rev. Respir. Dis. (1980), 122(5, Pt. 1), 808-9

CODEN: ARDSBL; ISSN: 0003-0805

DOCUMENT TYPE:

Journal; General Review

LANGUAGE:

English

A review with 11 refs. of evidence to indicate that S-carboxymethyl cysteine [638-23-3] is not purely a mucolytic agent but is a mucoregulator capable of normalizing the secretory disorders of the bronchial mucosa.

TΤ 638-23-3

RL: BIOL (Biological study)

(mucus secretion by bronchi response to)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 68 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1980:400482 HCAPLUS

DOCUMENT NUMBER:

93:482

TITLE:

The effect of mucolytic agents on the rheologic and

transport properties of canine tracheal mucus

AUTHOR(S):

Martin, Roberto; Litt, Mitchell; Marriott, Christopher Dep. Bioeng., Univ. Pennsylvania, Philadelphia, PA,

CORPORATE SOURCE: 19104, USA

Am. Rev. Respir. Dis. (1980), 121(3), 495-500

CODEN: ARDSBL; ISSN: 0003-0805

DOCUMENT TYPE:

Journal

LANGUAGE:

SOURCE:

English

AB The effect of several SH and other agents on the rheol. and mucociliary transport properties of a model secretion, reconstituted canine tracheal mucus, was investigated. The mucus was obtained via the canine tracheal pouch. Rheol. properties were detd. by microrheometry, and the ciliary transport rate was detd. using the frog palate technique. N-Acetyl cysteine [616-91-1] decreased the elastic modulus, leading to improved mucociliary transport at concns. such that the mucin did not ppt. S-Carboxymethyl cysteine [638-23-3] had no effect on either mucus properties or mucociliary transport rate, and its reported effectiveness in vivo must be due to some mechanism other than solubilization of mucin. Similar results were found with other blocked SH compds. Urea and KI did decrease mucus elasticity, but were harmful to cilia at the concns. needed.

TΥ 638-23-3

RL: BIOL (Biological study)

(tracheal mucus transport response to)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

HO₂C

L48 ANSWER 69 OF 81 HCAPLUS COPYRIGHT 2002 ACS 1981:435443 HCAPLUS

ACCESSION NUMBER:

95:35443 DOCUMENT NUMBER:

Tracheobronchial function in health and disease. TITLE:

Effect of mucolytic substances

Melville, G. Norris; Ismail, S.; Sealy, C. AUTHOR(S):

Physiol. Dep., Univ. West Indies, Kingston, Jamaica CORPORATE SOURCE:

Respiration (1980), 40(6), 329-36 SOURCE: CODEN: RESPBD; ISSN: 0025-7931

DOCUMENT TYPE: Journal English LANGUAGE:

The effect of mucolytic and expectorant substances on ciliary beat frequency, mucus transport velocity, and mucus prodn., was investigated in normal and bronchitic rats. N-Acetylcysteine [616-91-1] and S-carboxymethylcysteine [638-23-3] were mildly cilioexcitatory at low and ciliodepressive at higher concns. in both normal and bronchitic rats. A similar pattern was seen in mucus transport velocity. Bisolvon [611-75-6] enhanced all aspects of mucociliary activity in both groups of animals. Sobrepin [78006-54-9] was less effective than Bisolvon and more effective than Tachoquilin [9046-29-1]. Gelomyrtol [8002-55-9], Ozothine [8031-65-0] and prostaglandin El [745-65-3] were all cilioexcitatory in rats with bronchitis. Mucus transport velocity was similarly stimulated by both Gelomyrtol and Ozothine. Ammonium chloride and potassium iodide enhanced mucociliary activity in normal and bronchitic rats. All substances stimulated mucus prodn., however, the most potent was prostaglandin E1. The mechanisms for increased mucociliary activity involve inter alia the probable cleaving of disulfide bridges, decreased mucosal swelling, altered rheol. characteristics and stimulation of adenylate cyclase.

638-23-3 IT

RL: BIOL (Biological study) (ciliary beat frequency and mucus transport and mucus prodn. response

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$_{\mathrm{HO_{2}C}}$$
 s $_{\mathrm{NH_{2}}}^{\mathrm{R}}$ $_{\mathrm{CO_{2}H}}^{\mathrm{CO_{2}H}}$

L48 ANSWER 70 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1979:581454 HCAPLUS ACCESSION NUMBER:

91:181454 DOCUMENT NUMBER:

Therapeutic compositions with bacteriolytic and TITLE:

mucolytic action

Prugnaud, Robert Louis INVENTOR(S):

PATENT ASSIGNEE(S): Fr.

Fr. Demande, 18 pp. SOURCE:

CODEN: FRXXBL

Patent DOCUMENT TYPE:

LANGUAGE: French

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE FR 2398497 19790223 FR 1977-22715 A1 19770725

A mucolytic and bactericidal compn. contains ampicillin AB

[69-53-4] and S-carboxymethylcysteine [638-23-3].

IT 638-23-3

RL: BIOL (Biological study)

(bactericidal-mucolytic compn. contg. ampicillin and)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

L48 ANSWER 71 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1980:33658 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 92:33658

TITLE: Effects of S-carboxymethylcysteine on the absorption

of orally administered amoxicillin in rats

AUTHOR(S): Broccali, G.; Nusdeo, O.

Lab. Ric. Biomed., Ital. Soc. Farm., Trezzano, Italy CORPORATE SOURCE:

Riv. Farmacol. Ter. (1979), 10(2), 173-8 SOURCE:

CODEN: RVFTBB; ISSN: 0302-1750

DOCUMENT TYPE: Journal Italian

LANGUAGE:

GT

The absorption of intragastrically administered Na amoxicillin (I Na salt) AB [34642-77-8] (100 mg/kg) in rats was enhanced by the simultaneous administration of the mucolytic compd. S-carboxymethylcysteine [638-23-3] (30 mg/kg), as shown by increased I concns. in the serum, lungs, and inflammatory exudate.

IT 638-23-3

RL: BIOL (Biological study)

(amoxicillin absorption by digestive tract increase by)

RN 638-23-3 HCAPLUS

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Page 54 09/868106 Cook

CO2H R HO₂C NH2

L48 ANSWER 72 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1978:191491 HCAPLUS ACCESSION NUMBER:

88:191491 DOCUMENT NUMBER:

Sulfur-containing N-benzylamino acids TITLE:

Baille-Barrelle; Vigneron, Maurice; Lespagnol, Charles INVENTOR(S):

Laboratoires Boehringer Ingelheim, Fr. PATENT ASSIGNEE(S):

Ger. Offen., 27 pp. SOURCE: CODEN: GWXXBX

Patent DOCUMENT TYPE:

German LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 2628911	A1	19780112	DE 1976-2628911	19760628
FI 7701834	A	19771229	FI 1977-1834	19770610
FI 68613	В	19850628		·
FI 68613	Č	19851010		
AT 7704215	A	19780815	AT 1977-4215	19770615
AT 348983	В	19790312		
NO 7702237	A	19771229	NO 1977-2237	19770624
NO 144421	В	19810518		
NO 144421	C	19810826		
BE 856157	A1	19771227	BE 1977-178820	19770627
DK 7702854	A	19771229	DK 1977-2854	19770627
DK 146445	В	19831010		
DK 146445	Č	19840319		
SE 7707418	Ā	19771229	SE 1977-7418	19770627
SE 441093	В	19850909	•	
SE 441093	č	19851219		
NL 7707085	A	19771230	NL 1977-7085	19770627
JP 53031630	A2	19780325	JP 1977-76438	19770627
ES 460117	A1	19781001	ES 1977-460117	19770627
ZA 7703835	A	19790228	ZA 1977-3835	19770627
US 4185114	A	19800122	US 1977-810087	19770627
CH 629483	A	19820430	CH 1977-7873	19770627
FR 2357539	A1	19780203	FR 1977-19825	19770628
FR 2357539	B1	19821112		
GB 1565411	A	19800423	GB 1977-27068	19770628
GB 1565412	A	19800423	GB 1979-197	19770628
GB 1565413	A	19800423	GB 1979-198	19770628
AT 7803143	A	19781115	AT 1978-3143	19780502
AT 350523	В	19790611		
AT 7803144	Ā	19790115	AT 1978-3144	19780502
AT 351506	В	19790725		
AT 7803145	Ā	19790115	AT 1978-3145	19780502
AT 351507	В	19790725		
ES 470892	A1	19790201	ES 1978-470892	19780616
ES 470890	A1	19790201	ES 1978-470890	19780616
ORITY APPLN.		= -	DE 1976-2628911	19760628
WILL WILDI.	11,10.,		GB 1977-2628911	19760628
			AT 1977-4215	19770615
			GB 1977-27068	19770628

09/868106

R—CHNHCH
$$(CO_2R^3)$$
 $(CH_2)_nSR^4$

$$NHR^2$$

N-benzyl S-contg. amino acids I (R and R1 = H, halo; R2 = H, aliph, acyl; R3 = H, C1-4 alkyl; R4 = H, C1-3 alkyl, carboxy lower alkyl, acyl; n = 1, 2), their inorg. or org. acid addn. salts, their sulfonic acid salts, and, if R3 = H, their compds. with inorg. or org. bases or basic amino acids were prepd. as agents for the treatment of bronchial hypersecretions produced by irrations. Thus, methionine was treated with o-nitrobenzaldehyde to give the Schiff base which was reduced with NaBH4 to give o-O2NC6H4CH2-Met-OH (II). II was hydrogenated over Raney Ni to give 80% o-H2NC6H4CH2-Met-OH which was brominated by Br in HOAc at 60-65.degree. to give 27.5% I (R = R1 = Br, R2 = R3 = H, R4 = Me) (III). III at 468.7 mg/kg was used in guinea pigs against bronchial hypersecretions produced by inhalation of NH3 vapor.

Τ

IT 638-23-3

RL: RCT (Reactant)

(reaction of, with nitrobenzaldehyde)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 73 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1978:471708 HCAPLUS

DOCUMENT NUMBER: 89:71708

TITLE: The biological fate of vinylidene chloride in rats

AUTHOR(S): Jones, B. K.; Hathway, D. E.

CORPORATE SOURCE: Cent. Toxicol. Lab., Imp. Chem. Ind. Ltd., Alderley

Park/Cheshire, Engl.

SOURCE: Chem.-Biol. Interact. (1978), 20(1), 27-41

CODEN: CBINA8; ISSN: 0009-2797

DOCUMENT TYPE: Journal LANGUAGE: English

The main eliminative route for 14C-labeled vinylidene chloride (I) [75-35-4] after intragastric, i.v. or i.p. administration to rats was pulmonary; both unchanged I and I-related CO2 were excreted by that route and other I metabolites via the kidneys. Part of the urinary I was of biliary origin. Pulmonary elimination of I and CO2 and urinary excretion of I metabolites after an intragastric dose took 3 days, whereas >60 and 80% of a small i.v. dose were excreted unchanged within 5 min and 1 h after injection, resp. Biotransformation of I gave thiodiglycollic acid [123-93-3] and an N-acetyl-S-cysteinyl-acetyl deriv. as major urinary metabolites together with substantial amts. of chloroacetic acid [79-11-8], dithioglycollic acid [505-73-7], and thioglycollic acid [68-11-1]. It is probable that chloroacetic acid, which is a I metabolite per se, lies on a main metabolic pathway for I, since it affords several metabolites in common with I. Electrolysis of 1 mol. proportion of the

Page 56 09/868106 Cook

thiodiglycollate metabolite from I or chloroacetic acid gave 1 equiv. of CO2; this evidence is consistent with the transformation of I into chloroacetic acid by a mechanism involving the migration of one Cl atom and the loss of the other one. CO2 may be produced through the action of epoxide hydratase on 1,1-dichloroethylene oxide or by a minor oxidative pathway for chloroacetic acid. The N-acetyl-S-cysteinyl-acetyl deriv. is probably formed via the reaction of 1,1-dichloroethylene oxide and glutathione S-epoxide transferase.

638-23-3 ΙT

RL: FORM (Formation, nonpreparative)

(formation of, in vinylidene chloride metab.)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

$$R = CO_2H$$
 NH_2

L48 ANSWER 74 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1976:414144 HCAPLUS ACCESSION NUMBER:

85:14144 DOCUMENT NUMBER:

Soluble organic compounds containing sulfur

TITLE: Societe d'Etudes et Applications Chimiques, Fr. PATENT ASSIGNEE(S):

Belg., 11 pp. SOURCE:

CODEN: BEXXAL

Patent DOCUMENT TYPE: French LANGUAGE: FAMILY ACC. NUM. COUNT: 1 .

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
BE 827639 FR 2284319 ES 438100 NL 7508113 NL 162361	A1 A1 A1 A B	19750731 19760409 19770116 19760312 19791217	BE 1975-155156 FR 1974-30693 ES 1975-438100 NL 1975-8113	19750407 19740910 19750531 19750708
NL 162361 GB 1501138 CH 602735 JP 51054590 DE 2539863	C A A A2 A1	19800516 19780215 19780731 19760513 19760318	GB 1975-28961 CH 1975-9519 JP 1975-107947 DE 1975-2539863	19750709 19750721 19750905 19750908
ORITY APPLN. INFO.	:		FR 1974-30693	19740910

PRIO Five double salts consisting of a 7-theophylline acetic acid deriv., AB ethylinediamine [107-15-3], and a mercaptoamino acid were effective as respiratory agents. Thus, these salts inhibited histamine-induced broncho spasms, respiratory depression from either morphine or barbiturates, and caused a regression of mucopurulent obstructions from SO2 inhalation. These salts were prepared from solns. contg. equimolar portions of the starting compds.

TΨ 638-23-3

RL: RCT (Reactant)

(reaction of, with ethylenediamine and theophylline acetic acid)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

L48 ANSWER 75 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1975:453126 HCAPLUS

DOCUMENT NUMBER: 83:53126

TITLE: Bronchopulmonary uptake of sulfur from

> carboxymethylcysteine-35S and cysteine-35S in normal rats and rats exposed to sulfur dioxide. Effects of

prolonged administration of these two

sulfur-containing molecules on bronchopulmonary lesions and on pulmonary uptake of their sulfur

AUTHOR(S): Servin, A.; Garcet, S.; Vu Ngoc Huyen CORPORATE SOURCE: Cent. Rech., Lab. Joullie, Puteaux, Fr.

SOURCE: Bull. Physio-Pathol. Respir. (1974), 10(3), 315-29

.CODEN: BPPRA6

DOCUMENT TYPE: Journal French LANGUAGE:

AB The uptake of radioactivity by the lungs of normal rats after a single oral dose of 35S-labeled S-carboxymethylcysteine [638-23-3] was more tissue-specific than that after labeled cysteine [52-90-4], the tissue-to-plasma ratios of 35S being 2 and 1, resp. In rats with pulmonary lesions consequent to SO2 exposure (used as a model of chronic bronchitis), the tissue-to-plasma ratio of S-carboxymethylcysteine was 3 after a single dose, and 4 after a 5-week treatment (500 mg/kg/day, orally). In the latter case, an improvement in bronchial structure was obsd., esp. a decrease in mucosal goblet cell hyperplasia. This improvement was not seen in cysteine-treated animals, nor was the pulmonary specificity for cysteine altered by 5-week administration. These differences in distribution and pharmacol. activity can be partially explained by the fact that cysteine S was excreted in the urine as inorg. sulfates, whereas S-carboxymethylcysteine was excreted mainly unchanged.

IT 638-23-3

RL: BIOL (Biological study)

(bronchopulmonary damage from sulfur dioxide response to and lung metab. of)

638-23-3 HCAPLUS RN

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 76 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1975:11333 HCAPLUS

DOCUMENT NUMBER: 82:11333

TITLE: Action of chronic administration of

S-carboxymethylcysteine on pulmonary protein synthesis

disturbances induced by sulfur dioxide in rats

AUTHOR(S): Garcet, S.; Servin, A.; Vu Ngoc Huyen

CORPORATE SOURCE: Cent. Rech., Lab. Joullie, Puteaux, Fr.

SOURCE: C. R. Seances Soc. Biol. Ses Fil. (1974), 168(1), 43-6

CODEN: CRSBAW

DOCUMENT TYPE: Journal LANGUAGE: French

The increased protein formation obsd. in the bronchopulmonary region of rats with exptl. bronchitis induced by SO2 was returned to normal by prolonged oral administration of S-carboxymethylcysteine [638-23-3] (500 mg/kg/day).

ΙT 638-23-3

RL: BIOL (Biological study)

(protein formation response to, in bronchitis)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

S R CO₂H

L48 ANSWER 77 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1973:52507 HCAPLUS ACCESSION NUMBER:

78:52507 DOCUMENT NUMBER:

Tissue distribution of sulfur-35 as a function of time TITLE:

after oral administration of carboxymethylcysteine-

35S. Autoradiography and pharmacokinetics

Servin, A.; Garcet, S.; Vu Ngoc Huyen; Muller, P.; AUTHOR(S):

Cohen, Y.

Lab. Pharmacodyn., Fac. Pharm., Paris, Fr. CORPORATE SOURCE:

C. R. Soc. Biol. (1972), 166(4-5), 543-8 SOURCE:

CODEN: CRSBAW

DOCUMENT TYPE: Journal French LANGUAGE:

35S-labeled S-carboxymethylcysteine [638-23-3], a drug active on AB the bronchial mucosa, was very rapidly absorbed after oral administration to mice and was excreted mainly by the kidneys. Autoradiog. and quant. detns. showed the 35S to have particular affinity for the lungs, which took up radioactivity more slowly (max. after 3 hr) but retained it much longer than did the other organs. The activity reached its highest abs. level in the pancreas, whereas it was practically nil in the brain and

heart. ΙT 638-23-3

RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (metabolism of)

638-23-3 HCAPLUS RN

L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

HO₂C S R CO₂H

L48 ANSWER 78 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1971:409880 HCAPLUS ACCESSION NUMBER:

75:9880 DOCUMENT NUMBER:

Mucolytic formulations containing TITLE:

S-(carboxymethyl)cysteine

Joullie, Maurice; Vu-Ngoc-Huyen; Maillard, Gabriel; INVENTOR(S):

Lakah, Lucien; Muller, Pierre

PATENT ASSIGNEE(S): Recherches Pharmaceutiques et Scientifiques

SOURCE: Ger. Offen., 11 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent German LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 2006486	A	19710422	DE 1970-2006486	19700213
FR 2068398	A6	19710827	FR 1969-34139	19691007
FR 2068398	В2	19730112		
GB 1272881	A	19720503	GB 1970-1272881	19700213
ZA 7006363	A	19710527	ZA 1970-6363	19700917
US 3891749	А	19750624	US 1973-389325	19730817
US 29256	E	19770607	US 1976-692031	19760602
PRIORITY APPLN. I	NFO.:		FR 1969-34139	19691007
			US 1970-10999	19700212
			US 1973-389325	19730817

AΒ Formulations of the title compd. (I), optionally in admixt. with antibiotics, antihistaminics, corticosteroids, or bronchodilators, for oral administration were reported. A typical compn. of a capsule was I 50, lactose 40, silica gel 5, and Mg stearate 5 mg.

638-23-3 IT

RL: BIOL (Biological study)

(mucolytic agent)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L48 ANSWER 79 OF 81 HCAPLUS COPYRIGHT 2002 ACS

1967:420418 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 67:20418

TITLE: Effect of mucolytic compounds on the experimental

intrabronchial mucus retention in rats

AUTHOR(S): Quevauviller, Andre; Huyen-Vu-Ngoc; Garcet, Suzanne;

Lakah, Lucien

CORPORATE SOURCE: Fac. Pharm., Paris, Fr.

SOURCE:

Therapie (1967), 22(2), 485-93

CODEN: THERAP

DOCUMENT TYPE: Journal LANGUAGE: French

Bronchial hypersecretion was induced in female Wistar rats by exposure to SO2 for a total of 110 hrs. over a 16-day period. Some of the rats were treated orally simultaneously with S-carboxymethyl-cysteine (I) (8 g./kg./animal). Twenty-four hrs. after treatment, the rats were sacrificed and the bronchial system observed macroscopically and histol. This in vivo method for detg. mucolytic activity by measuring the retention of intrabronchial mucus in the rat gave evidence of mucolytic activity by I. I had lytic power similar to that of aerosols contq. reducing agents with SH groups.

IT 2387-59-9

> RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mucolytic activity of)

RN 2387-59-9 HCAPLUS

L48 ANSWER 80 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1969:9718 HCAPLUS

DOCUMENT NUMBER: 70:9718

TITLE: The nitrogen pool of animal tissues. Ox liver and

bile. Ox kidney and lung

AUTHOR(S): Azumi, Tsukasa

CORPORATE SOURCE: Med. Sch., Okayama Univ., Okayama, Japan SOURCE: Acta Med. Okayama (1967), 21(6), 321-6

CODEN: AMOKAG

DOCUMENT TYPE: Journal LANGUAGE: English

AB The compns. of N pools of ox liver, bladder bile, kidney, and lung were analyzed with special emphasis on the minor components, and some distinctive features of these tissues were described.

S-(1,2-Dicarboxyethyl-L-cysteine and 3-(carboxymethyl)-L-cysteine were found in ox liver and kidney. Liver was low in free arginine and lysine, but high in ornithine, ethanolamine, and glutathione. Glycine was predominant only in ox bile. All amino acids were present in moderate amts. in kidney, but the glutathione content was low. The concns. of arginine and lysine were relatively high in lung.

IT 638-23-3

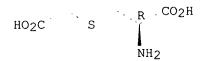
RL: BIOL (Biological study)

(in organs)

RN 638-23-3 HCAPLUS

CN L-Cysteine, S-(carboxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L48 ANSWER 81 OF 81 HCAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1967:114378 HCAPLUS

DOCUMENT NUMBER: 66:114378

TITLE: Experimental hypersecretion of bronchial mucus in the

rat. II. Application to the study of

S-(carboxymethyl)cysteine

AUTHOR(S): Huyen-Vu-Ngoc; Garcet, Suzanne; Lakah, Lucien

CORPORATE SOURCE: Serv. Rech., Lab. Joullie, Puteaux, Fr.

SOURCE: C. R. Seances Soc. Biol. Ses Fil. (1966), 16(10),

1849-51

CODEN: CRSBAW

DOCUMENT TYPE: Journal LANGUAGE: French

AB Rats were made to breathe air contg. 0.03% of SO2 2-5 hrs./day, 5 days/week, for 16 days. Half or them were given oral doses of S-(carboxymethyl)cysteine, 500 mg./kg./day, during the exptl. period. Without the drug treatment 71% of the rats exposed to SO2 developed severe obstruction of the bronchi by purulent mucus but only 17% of those given the drug showed this effect. The drug apparently has considerable mucolytic activity.

IT 2387-59-9

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (mucolytic activity of)

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ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS
L49
     25390-17-4 REGISTRY
RN
CN
     Cysteine, S-(carboxymethyl) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Alanine, 3-[(carboxymethyl)thio]-, DL- (8CI)
CN
     DL-Cysteine, S-(carboxymethyl)-
CN
OTHER NAMES:
     5-Amino-3-thiadihexanoic acid
CN
CN
     DL-3-(Carboxymethylthio)alanine
     S-(Carboxymethyl)-(RS)-cysteine
CN
CN
     S-(Carboxymethyl)-DL-cysteine
CN
     S-(Carboxymethyl)cysteine
FS
     3D CONCORD
DR
     2387-59-9
MF
     C5 H9 N O4 S
     COM
CI
                 ANABSTR, BEILSTEIN*, BIOSIS, CA, CANCERLIT, CAOLD, CAPLUS,
LC
     STN Files:
       CASREACT, CHEMCATS, CHEMLIST, CSCHEM, IPA, MEDLINE, NIOSHTIC, RTECS*,
       TOXCENTER, USAN, USPATFULL
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     Other Sources:
                      EINECS**
         (**Enter CHEMLIST File for up-to-date regulatory information)
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              20 REFERENCES IN FILE CAPLUS (1967 TO DATE)
L49
     ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS
RN
     638-23-3 REGISTRY
     L-Cysteine, S-(carboxymethyl) - (9CI) (CA INDEX NAME)
CN
OTHER CA INDEX NAMES:
     Alanine, 3-[(carboxymethyl)thio]-, L- (6CI, 8CI)
OTHER NAMES:
     (L)-2-Amino-3-(carboxymethylthio)propionic acid
CN
CN
     (R) -S-(Carboxymethyl) cysteine
CN
     3-[(Carboxymethyl)thio]-L-alanine
     Bronchokod
CN
CN
     Carbocisteine
CN
     Carbocysteine
CN
     DF 1794Y
CN
    L-(Carboxymethyl)cysteine
CN
     LJ 206
     Muciclar
CN
CN
     Mucodyne
CN
     Mucopront
CN
     Rhinathiol
     Rhinatiol
CN
CN
     Rinatiol
CN
     S-(Carboxymethyl)-(R)-cysteine
CN
     S-(Carboxymethyl)-L-cysteine
CN
     S-Carboxylmethyl-L-cysteine
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CN

Thiodril

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AR 2387-59-9

FS STEREOSEARCH

DR 11139-64-3

MF C5 H9 N O4 S

CI COM

LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DRUGU, EMBASE, HODOC*, IFICDB, IFIPAT, IFIUDB, MRCK*, NIOSHTIC, PHAR, PHARMASEARCH, PROMT, RTECS*, TOXCENTER, ULIDAT, USPATFULL, VETU

(*File contains numerically searchable property data)
Other Sources: EINECS**, NDSL**, TSCA**, WHO
 (**Enter CHEMLIST File for up-to-date regulatory information)

Absolute stereochemistry.

$$HO_2C$$
 S R CO_2H NH_2

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

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RN 2387-59-9 HCAPLUS

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